M. musculus 107390 Sequence 4

BD057272 Gene enco
AB057272 Gene enco
CS125505 Sequence
CS126190 Sequence
CS126190 Sequence
AJ2561 Musm muscu
A22561 Musmcsulus
AR09372 Sequence
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AX080054 Sequence
A7118 Sequence
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A700058 Musmuscu
BC10835 Mus muscu
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BC108375 Mus muscu
BC001889 Mus muscu
BC01853 Mus muscu
BC01853 Mus muscu
BC01858 Mus muscu
BC01858 Mus muscu
BC01889 Humanized a
BD022360 Multi-fun

protein

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Glaichenhaus, N. and Malherbe, L.
Recombinant proteins and molecular complexes derived therefrom, analogous to molecules involved in immune responses
Patent: WO 0109194-A 1 08-FEB-2001,
CENTRE NATIONAL DE LA RECHERCHE SCIENTIFIQUE (CNRS) (FR)
Location/Qualifiers

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CS125905
CS126190
CS138860
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       AX081280 Sequence
BD137962 Monovalen
AX080953 Sequence
                                                May 31, 2006, 22:52:13 ; Search time 9316.63 Seconds (without alignments) 5096.366 Million cell updates/sec
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                                                                                           MPCSRALILGVLALNTMLSL.......VHEGLHNHHTTKSFSRTPGK 495
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Description
       GenCore version 5.1.8
Copyright (c) 1993 - 2006 Biocceleration Ltd.
                                    nucleic search, using frame_plus_p2n model
                                                                                                                                                                 hits satisfying chosen parameters:
                                                                                                                                                    6366136 segs, 31973710525 residues
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Maximum Match 100%
Listing first 45 summaries
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PAT 27-FEB-2001

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CQ806531 S CQ806532 S AR577144 S AX280226 S AR650819 S

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BD137962 1446 bp DNA linear PAT 18-SEP-2002
Nonovalent MHC-binding domain fused proteins and conjugates,
polyvalent MHC-binding domain fused proteins and conjugates,
polymer MHC-binding domain fused proteins and conjugates,
utilization therefor.
BD137962 BD137962. GI:22232907
SPINFACT. GI:22232907
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                   GluargThr11eSerly8ProLy8GlySerValArgAlaProGlnValTyrValLeuPro
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PRESIDENT AND FELLOWS OF HARVARD COLLEGE
OS Artificial Sequence
PN JP 200250442-A/7
PD 12-FEB-2002
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Flt-3 ligand-encoding polynucleotide ав a polynucleotide-based
vaccine enhancer
                                      ThrThr ------AlaProSerAlaGlnLeuGluLysGluLeuGlnAlaLeuGlu
                                                            610 TTAACTGATACACTCCAAGCGGAGACAGATCAACTTGAAGACGAGAAGTCTGCGTTGCAG
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19-FEB-1999 JP 2000532537
19-FEB-1998 US 60/075351
KAI W WUCHERPFENNIG, JACK L STROMINGER
C12N15/09, A61K35/14, A61K47/48, C07K14/705, C07K16/00, C07K19/00,
C12Q1/02;
GQ1N33/53,C12N15/00
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BRA*0101 extracellular domain
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Location/Qualifiers
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Immunogenic recombinant antibody
Patent: WO 20040916555-A 1 28-OCT-2004;
Igeneon Krebs-Immuntherapie Forschungs- und Entwicklungs- AG
Location/Qualifiers
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unclassified sequences.

1 (bases 1 to 1581)
Boulain,J. and bucancel,F.
BOULBIAIN,J. and bucancel,F.
APPLICATIONS AS AN AGENT IN DIAGNOSTIC AND IN THE THERAPEUTIC FIELD OR AS A REAGENT RELEVANT IN MEDICAL APPROACHES
PATENT: EP 0556111-A 1 18-AUG-1993;
COMMISSARIAT ENERGIE ATOMIQUE (FR)
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/ Droduct = Timunoglobulin, Heavy Chain"
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/ Drotein id="CAB58933.1"
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/organism="unidentified"
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Bukaryota; Metazoa; Chordata; Craniata; Glires; Rodentia;
Sciurognathi; Muroidea; Muridae; Murinae; Mus.
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S Ducancel, F.F.D.
Direct Submission
L Submitted (09-FEB-1993) F.F.D. Ducancel, C E A, Die de Saclay, 91191 Gif Yvette, Cedex, FRANCE
Location/Qualifiers
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                                            IGLUHIS-TrpGlyLeuGluGluProValLeuLysHisTrpGluProGluIleProAlaP
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Mismatches:
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Gaps:
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PAT 27-AUG-2002

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1157 AAACAGAGCTAAACTACAAGAACACTGAACCAGTCCTGGACTCTGATGGTTCTTACTTCA 1216
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C12N5/00
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HIDEO OKAWA, MASANOBU NAKATA, YOJIRO YUASA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    synthetic construct
other sequences; artificial sequences.
1 (Dases 1 to 1570)
Okawa, H., Nakata, M. and Yuasa, Y.
Gene encoding antimalathion monoclonal antimalathion monoclonal antimalathion monoclonal antimalathion monoclonal patent: JP 2001275682-A 9 09-CCT-2001;
KANKYO MENEKI GIJUTSU KENKYUSYO KK
PN JP 2001275682-A/9
PD 09-CCT-2001
PP 31-MAR-2000 JP 2000098323
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Beavers, L.S., Bumol, T.F., Gadski, R.A. and Weigel, B.J.
Novel recombinant and chimeric antibodies directed against a human adenocarcinoma antigen
Patent: EP 0338767-A2 4 25-OCT-1989;
Location/Qualifiers
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   1154 TCCACCAGAAGAAGAGATGACTAAGAAACAGGTCACTCTGACCTGCATGGTCACAGACTT 1213
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Mammalia, Eutheria, Euarchontoglires, Glires, Rodentia,
Sciurognathi, Muroidea, Muridae, Murinae, Mus.
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Location/Qualifiers
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/db_xref="taxon:10090"
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/product="immunoglobulin gamma-2a heavy chain"
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/note="Balb/c IgG2a hinge"
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Glaxo Group Limited (GB)
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/organism="Mus musculus"
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/db_xref="taxon:10090"
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                                              -----GlnTyrThrHisGluPheAspGlyAspGluLeuPheTyrValAspLeuAspLys
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Mus musculus
Bukaryotas, Metazoa, Chordata; Craniata; Vertebrata; Buteleostomi;
Mammalia; Butheria; Buarchontoglires; Glires; Rodentia;
Sciurognathi; Muroidea; Muridae; Murinae; Mus.
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                                             YSerTyrPheMetTyrSerLysLeuArgValGluLysLysBanTrpValGluArgAsnSe
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1143 GGTCACTCTGACTGCTCACAGACTTCATGCCTGAAGACATTTACGTGGAGTGGAC
                                                                                                                                                                       rasnasnGlyLysthrGluLeudsnTyrLysasnThrGluProValLeudspSeraspGl
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SerLysSerValThrAspGlyValTyrGluThrSerPheLeuValAsnArgAspHisSer 171
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    ->990
    /gene="IGHG2A"
    /product="immunoglobulin heavy chain constant region gamma 2a"

990 bp mRNA linear ROD 09-PEB-2001 Mus musculus partial mRNA for immunoglobulin heavy chain constant region gamma2a (IGHG2A gene).
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    product="immunoglobulin heavy chain constant region gamma
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Submitted (18-SEP-2000) McLean G.R., Cell Biology, Albert Einstein
College of Medicine, 1300 Morris Park Avenue, Bronx, New York
10461, USA
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                                                                                                                                                                                                                                 McLean,G.R., Nakouzi,A., Casadevall,A. and Green,N.S.
Human and murine immunoglobulin expression vector cassettes
Mol. Immunol. 37 (14), 837-845 (2000)
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                                                                                                                                                                  Bukaryota, Metazoa, Chordata, Craniata, Vertebrata, Butu
Mammalia, Butheria, Buarchontoglires, Glires, Rodentia,
Sciurognathi, Muroidea, Muridae, Murinae, Mus.
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Matches:
Conservative:
Mismatches:
Indels:
Gaps:
                                                                                                              constant region; gamma 2a; IGHG2A gene;
Mus musculus (house mouse)
Mus musculus
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/organism="Mus musculus"
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/db_xref="taxon:10090"
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Mammalia, Eutheria, Euarchontoglires, Glires, Rodentia,
Sciurognathi, Muroidea, Muridae, Murinae, Mus.
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Sikorav, J.L., Auffray, C. and Rougeon, F.
Structure of the constant and 3' untranslated regions of Balb/c gamma 2a heavy chain messenger RNA
Nucleic Acids Res. 8 (14), 3143-3155 (1980)
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   1mmunoglobulin
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Matches:
Conservative:
Mismatches:
Indels:
complementary DNA; gamma-immunoglobulin;
Mus musculus (house mouse)
Mus musculus
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/organism="Mus musculus"
/mol_type="mRNA"
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/product="messenger RNA"
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68.5%
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Best Local Similarity:
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nLeuGluTrpGluLeuGlnAlaLeuGluLysGluLeuAlaGlnAlaAlaSerGluProAr

245

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ROD 17-NOV-2004

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Direct Submission
Submitted (08-JAN-2002) Institute of Zoology, Academia Sinica, 128
Academia Road, NanKang, Taipei 11529, Taiwan, ROC
Location/Qualifiers
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Matches:
Conservative:
Mismatches:
Indels:
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/organism="Mus musculus"
/mol type="mRNA"
/dbrain="BALB/c"
/db xref="taxon:10090"
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Mus musculus
Bukaryota, Metazoa; Chordata; Craniata; Vertebrata; Buteleostomi;
Bukaryota, Metazoa; Chordata; Craniata; Glires; Rodentia;
Bammalia; Butheria; Buarchontoglires; Glires; Rodentia;
Sciurognathi; Muroidea; Muridae; Murinae; Mus.
1 (bases I to 1407)
Lai, X. S., John, J. A. C., Guo, I. C., Chen, S. C., Fang, K. and Chang, C. Y.
In vitro efficiency of intra- and extracellular immunization with
mouse anti-YGNNV antibody against yellow grouper nervous necrosis
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             900 GGTGGAAAGAAATAGCTACTCCTGTTCAGTGGTCCACGAGGGTCTGCACAATCACCACAC 959
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Lai, Y.-S., John, C., Guo, I.-C., Chen, S.-C., Fang, K. and Chang, C.-Y.
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                                                            ollevalThrCysValValValAspValSerGluAspAspRroAspValGlnIleSerTr
                                                                                                                                                                                                                                        480 GTTTGTGAACAACGTGGAAGTACACACAGCTCAGACACAAACCCATAGAGAGTTACAA
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                                                                                                                      ACCATCCGTCTTCATCTTCCCTCCAAAGATCAAGGATGTACTCATGATCTCCCTGAGGCC
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                                             gGlyProThrIleLysProCysProProCysLysCysProAlaProAsnLeuLeuGlyGl
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AUTHORS
TITLE
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762	TAACCTCTTGGGTGGACCATCCGTCTTCATCTTCCCTCCAAAGATCAAGGATGTACTCAT 821	
300	tleserLeuserProllevalThrCygvalvalvalAgpvalserGluAgpAspProAs 320 	
320	pValGinileSerTrpPheValAsnAsnValGiuValHisThrAlaGinThrGinThrHi 340 	
340	sarggluaspTyrasnSerThrLeuargValValSerAlaLeuBrolleGlnHisGlnAs 360 	
360	DTrpmetSerGlyLysGluPhelysCysLysValasnAsnLysAspLeuProAlaPro11 380 	
380	eGluargThrIleSerLysProLysGlySerValArgAlaProGlnValTyrValLeuPr 400 	
400	OProProGluGluGluMetThrLysLysGlnValThrLeuThrCysMetValThrAspPh 420 	
420	emet Proglua apilety rvalglut rpthrasna angly Lysthrglubeua antyrly 440 	
440	BABIThrGluProValleuAspSerAspGlySerTyrPheMetTyrSerLysLeuArgVa 460 	
1302	GlulyslysasnfrpValGlubrgasnSerTyrSerCysSerValValHisGluGlyle 480 	
480	uHibarnHibHibThrLysSerPheSerargThrProGlylys 495 	

sch completed: June 1, 2006, 03:20:55 time: 9336.63 secs

Adw4282 DR2 IgG f Ab199027 IAS MBP 1 Ab199031 MBP 90-10 Ab199032 MBP 1-14 Ab19010 MBP 1-14

dneuce:

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Recombinant protein; alpha chain; beta chain; MHC; immunoglobulin; major histocompatibility complex; Pc region; antigen; T lymphocyte; immunostimulant; vaccine; infection; tumour; ss.
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AEE21946
ADS31748
ADS92750
AD007566
                 ADW44282
ABI99027
ABI99033
ABI99029
ABI99032
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AEC20762
ADL15084
ADV26108
ADM71834
AAQ12637
AAQ10537
AAQ1053
AAR30341
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Aat62850 Mouse sol Aaz51300 Murine im

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(CNRS ) CNRS CENT NAT RECH SCI.

Glaichenhaus N, Malherbe L;

WPI; 2001-182944/18. P-PSDB; AAB67480. New soluble recombinant protein, useful e.g. as immunostimulant, comprises dimeric major histocompatibility complex molecule fused to immunoglobulin Fc region.

Example 1; Page 31-33; 43pp; French.

The specification describes soluble recombinant proteins that comprise at least a dimer formed from the alpha and beta-chains of MHC (major histocompatibility complex) Class I and II molecules in which at least one chain has, attached to its C-terminus, at least part of the FC region of an immunoglobulin. The recombinant proteins, when linked to an antigent partic peptide, are used to count and/or purify antigen-reactive T lymphocytes and to characterize their phenotype, e.g. in preclinical evaluation of vaccines. They are also used as immunostimulants, particularly for vaccine development (against infections and tumours), to count and determine phenotype of autoreactive T cells in subjects with, or at risk of developing, autoimmune diseases, e.g. for staging or evaluating treatments, and (to purify and/or enrich Ag-reactive T cells from cell cultures or patient samples, for use in subsequent curative or preventative cellular therapy. The present sequence encodes a recombinant protein of the invention, comprising an alpha chain of MHC molecules

Sequence 1484 BP; 414 A; 394 C; 362 G; 314 T; 0 U; 0 Other;

	1484	494	0	0	0	0
	Length:		Conservative:	Mismatches:	Indels:	Gaps:
	1.06e-218	2655.00	100.0%	100.0%	99.8	ις.
gnment Scores:	d. No.:	ıre:	cent Similarity:	t Local Similarity:	ry Match:	1

10-048-116B-2 (1-495) x AAF55098 (1-1484)

1	Met ProCysSerArgAlaLeuIleLeuGlyValLeuAlaLeuAsnThrMetLeuSerLeu	20
1	ATGCCGTGCAGCAGCACTCTGATTCTGGGGGTCCTCGCCCTGAACACCATGCTCAGCTC	09
21		40
61		120
41		09
121	CAGTCTCCTGGAGACATTGGCCAGTACACACATGAATTTGATGGTGATGATCTTTTAT	180
61		80
181	GTGGACTTGGATAAGAAAACTGTCTGGAGGCTTCCTGAGTTTGGCCAATTGATACTC	240
81		100
241	TTTGAGCCCCAAGGTGGTGTTGTTTTTTTTTTTTTTTTT	300
101		120
301	ACTAAGAGGTCAAATTTCACCCCAGCTACCAATGAGGCTCCTCAAGCGACTGTTTCCCCC	360
121	LysSerProValLeuLeuGlyGlnProAsnThrLeuIleCysPheValAspAsnIlePhe	140
361		420
141	ProProValileAsnileThrTrpLeuArgAsnSerLysSerValThrAspGlyValTyr	160
421		480

ABI99041 standard; cDNA; 1676 BP.

y du	161	161 GluthrSerPheLeuValAsnArgAspHisSerPheHisLysLeuSerTyrLeuThrPhe 180 	0 0
	181		0
දු දි	541	ATCCCTTCTGATGATGACATTTATGACTGCAAGGTGGAGCACTGGGGCCTGGGGCCCTGGAGGAGCCG	
6 G	601		
ò	221	31yGlyGlySerThrThrAlaProSerAlaGlnLeuGluLysGluLeuGlnAlaLeuGlu 240	0
qa	661	GETGGAGGATCCACTACAGCTCCATCAGCTCGAAAAAGAGCTCCAGGCCCTGGAG 720	0
È	241	VysGluAsnAlaGlnLeuGluTrpGluLeuGlnAlaLeuGluLysGluLeuAlaGlnAla 260	0
qq	721	aaggaaaatgcacagctggaatgggagttgcaagcactggaaaaggaactggctcaggca 780	0
ò 5	261	#1aSerGluProArgGlyProThrIIeLyBProCyBProProCygLyGCyBProAlaPro 280 #1	
ò	281	AsnLeuLeuGlyGlyProSerValPhellePheProProLysIleLysAapValLeuMet 300	
qq	841		0
č	301		0
qq	106	ATCTCCCTGAGCCCCATAGTCACATGTGTGGTGGTGGATGTGAGCGAGGATGACCCAGAT 960	0
ò	321		0
qq	961	GTCCAGATCAGCTGGTTTGTGAACAACGTGGAAGTACACACAC	20
č	341	ArgGluAspTyrAsnSerThrLeuArgValValSerAlaLeuProlleGlnHisGlnAsp 360	0
Db 1	021	adadaddartracaacadtacreeeggreereggeeeereeeeareeeareeeareagae 1080	89
ζo	361		
ъ 1	081		40
٠ ک	381	GluargThrIleSerLysProLysGlySerValArgAlaProGlnValTyrValLeuPro 400	0 6
-	141	GAGAGAACCATCTCAAAACCCAAAGGGTCAGTAAGAGCTCCACAGGTATATGTCTTGCCT 1200	3 (
	401	ProproglugludumetThrLysLysGlnValThrLeuThrCysMetValThrAspPhe 420	0
П	201	ccaccadaadaadadardadaaaadadacacacrccadcrgcardgrcacadacrrc 1260	9
	421		
н	261		20
8	441		
-	321		80
	461	GluLysLysAsnTrpValGluArgAsnSerTyrSerCysSerValValHisGluGlyLeu 480	0
	381		40
8 8	481	HisaanHisHisThrThrLysSerPheSerArgThrProGly 494 	
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RESULT 2 ABI99041	5	openderd, chwb. 1676 RD	

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25-FEB-2002 (first entry)

Murine pCB223 coding sequence.

Mouse; MHC; major histocompatibility complex; MHC class II; multimer; single chain; immunosuppressive; antidiabetic; antidiamatory; antianaemic; anticheumatorid; antiarthritic; neuroprotective; vaccine; autoimmune disease; insulin dependent diabetes; multiple sclerosis; myasthemia gravis; permicious anaemia; autoimmune encephalomyelitis; rheumatoid arthritis; systemic lupus erythematosus; ss.

Mus sp.
Synthetic.

WO200170245-Al.

27-SEP-2001.

22-MAR-2000; 2000US-0191274P.

15-MAY-2000; 2000US-0264003P.

(CORI-) CORIXA CORP.

(CORI-) CORIXA CORP.

Carter D, Zhu S, Arimilli S, Wang A;

WPI; 2001-616371/71.
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Multimeric complex for treating autoimmune diseases, comprises first and second single chain MHC class II molecules, each comprising alphal and betal domain linked through amino acid linker and multimerization domain.

P-PSDB; ABB56471,

Disclosure; Page 115; 147pp; English.

The invention relates to a multimeric complex comprising a first recombinant single chain major histocompatibility complex (MHC) class II molecule and a second recombinant single chain MHC class II molecule, each comprising an alphal domain and a betal domain inked through an amino acid linker and a multimerisation domain. The first and the second multimeric complex, it useful for treating autoimmune diseases. It is useful for treating insulin dependent diabetes, multiple sclerosis, myasthenia gravis, pernicious anaemia, autoimmune encephalomyelitis (EAB), rheumatoid arthritis and systemic lupus erythematosus. The present sequence encodes a single chain MHC class II molecule of the invention

Sequence 1676 BP; 438 A; 470 C; 407 G; 361 T; 0 U; 0 Other;

	1676	432	13	39	64	2
	Length:	Matches:	Conservative:	Mismatches:	Indels:	Gaps:
	1.68e-183	2247.00	81.2%	78.8%	84.5%	4
ignment Scores:	3d. No.:	ore:	rcent Similarity:	st Local Similarity:	ery Match:	

-10-048-116B-2 (1-495) x ABI99041 (1-1676)

Gly 44     GGA 156	Asp 64
rThrValTyrGlnSerPrc 	45 AsplleGlyGlnTyrThrHisGluPheAspGlyAspGluLeuPheTyrValAspLeuAsp 64
ilglyPheTyrGlyThi        aggcgrcratggtac	uPheAspGlyAspGlt
leGluAlaAspHisVa                   TTGAGGCCGACCACGT	lyGlnTyrThrHisGl
25 AspAspI         97 GACGACA	45 AspileG
	25 ABPABDIleGluAlaABPHisValGlyPheTyrGlyThrThrValTyrGlnSerProGly 44

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CTCCGGGTGGTCAGTGCCCTCCCCATCCAGCACCAGGACTGGATGAGTGGCAAGGAGTTC
                                                                                                                               AAAGGGTCAGTAAGAGCTCCCACAGTATATGTCTTGCCTCCACCAGAAGAAGAAGATGACT
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Major histocompatibility complex class II; MHC class II; human; fusion protein; HLA-RR2; DRA*0101; binding domain; Fos; dimerisation domain; IgG; allergy; autoimmune disease; vaccine; multiple sclerosis; therapy; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                - comprising a MHC Class II or an immunoglobulin region
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Mus musculus.
Chimeric.
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17-AUG-1998
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This nucleotide sequences codes for a bivalent DR2 fusion protein obtained by fusion of the Fc portion of 1gG2a to the 3' end of a DR-alpha -Fos cDNA construct (see AAV16866). The Fc portion was amplified by RT-

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489
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PCR from mouse hybridoma L243. The PCR product was then fused in frame with the DR-alpha-Fos construct by overlapping PCR. The DR2-IgG fusion was expressed in the Drosophila Schneider cell system. The invention relates to new soluble monovalent and multivalent Class II MHC fusion proteins comprising a MHC class II binding domain and a dimerisation domain or an immunoglobulin region that can be used for the treatment of allergic and autchimume diseases (e.g. multiple sclerosis), for tolerising a subject to foreign tissue before or after organ or tissue transplantation, or for vaccination against pathogens. (Updated on 17-OCT -2003 to standardise OS field)
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Matches:
Conservative:
Mismatches:
Indels:
Gaps:
                                                                                                                                      356 G; 301
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83.2%
76.5%
                                                                                                                                        BP; 414 A; 375
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Best Local Similarity:
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Pred. No.:
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This nucleotide sequence codes for a divalent HLA-DR2 MHC binding domain fusion protein (see AAY31654) comprising an alpha-mating factor secretion signal, the extracellular domain of the HLA-DR2 alpha chain (residues 1-191 of DRA+0101), a 7-amino acid linker, the 40-amino acid leucine zipper dimerization domain of Fos, and the FC portion of IgG2a. The DR-alpha-FC chain corresponds to an antibody heavy chain. The invention provides new monovalent, multivalent and multimaric MHC Class II binding domain fusion proteins and conjugates comprising at least a binding domain of an MHC class II alpha or beta chain and a dimerization domain, especially a Fos or Jun leucine zipper domain. The MHC fusion proteins and conjugates can be used: for detecting and isolating T cells having a defined MHC/peptide complex (claimed); to stimulate or activate T cols a defined MHC/peptide complex (claimed); to stimulate or activate T cols a defined MHC/peptide complex (claimed); for selective to a defined MHC/peptide complex (claimed); to treat calergic and autoimmune diseases, e.g. multiple sclerosis, rheumatoid architis, pemphigus vulgaris, and systemic lupus erythematosus; and to prevent organ or tissue transplant rejection. The DR2-IgG design was chosen to increase the affinity for the T cell receptor by increasing valency, and to attach an effector domain, the Fc region of 1962a.

Complement fixation may result in the lysis of target T cells following the prince of target T cells following that therefore be useful for the selective depletion of autoaggressive T
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              26 AspileGluAlaAspHisValGlyPheTyrGlyThrThrValTyrGlnSerProGlyAsp
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                                                                                                                                                                                                                                                                                                                                                                                                               New HMC Class II binding domain fusion proteins and conjugates - used for, e.g. treating allergic and autoimmune diseases or detecting, isolating, activating or killing specific T cells.
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Conservative:
Mismatches:
Indels:
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                                         "DR2-Fos-Fc"
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  GCAGCATCTGAGCCCAGAGGGCCCACAATCAAGCCCTGTCCTCCATGCAAATGCCCAGCA 789
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                                                                                                                                                                                                                   AspvalGlnIleSexTxpPhevalAsnAsnValGluValHisThrAlaGlnThrGlnThr
                                                                                                                                                                                                                                                                                                        340 HisargGluaspTyrasnSerThrLeuargValValSerAlaLeuProIleGlnHisGln
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                                                                              790 CCTAACCTCTTGGGTGGACCATCCGTCTTCATCTTCCCTCCAAAGATCAAGGATGTACTC
                                                                                                                                300 MetileSerLeuSerProlleValThrCysValValValAspValSerGluAspAspPro
                                                                                                                                                       ATGATCTCCCCTGAGCCCCCATAGTCACATGTGGTGGTGGATGTGAGCGAGGATGATGACCCA
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/note= "alpha-mating factor secretion signal"
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Chimeric.
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AAGGAGACGGTCTTGGAGGATTTTGGACGATTTTGCCAGCTTTTGAGGCTCAAGGT
                       GlyLeuGlnAsnIleAlaAlaGluLysHisAsnLeuGlyIleLeuThrLysArgSerAsn
                                                                                                                                                  AlaAlaSerGluProArgGlyProThrIleLysProCysProCysLysCysProAla
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                                                                                       479
440 LybabithrGluprovalleuabpSerabpGlySerTyrPheMetTyrSerLybLeuarg
                                                                                                              ValGluLysLysAsnTrpValGluArgAsnSerTyrSerCysSerValValH1sGluGly
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/note= "DRA*0101extracellular domain"
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/*tag= b
/product= "DR2-IgG fusion
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/note= "Linker sequence"
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97WO-US014503.
98US-0075351P.
99US-00248964.
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                                                                                                                                                                                                                                                                                                                                                                                                     (first entry
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P-PSDB; ADW44283.
                                                                                                                                                                                                                                                                                                               standard;
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19-FEB-1998;
12-FEB-1999;
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Novel class II major histocompatibility complex (MHC) fusion protein having MHC class II binding domain of MHC class II alpha chain, and dimerization domain, useful for treating pemphigus vulgaris, rheumatoid arthritie.

Example; SEQ ID NO 11; 55pp; English.

Bystemic The present invention relates to the class II major histocompatibility complex (MHC) fusion protein having MHC class II binding domain of MHC class II alpha chain and a dimerization domain. The invention is useful in adoptive immunotherapy and tolerizing against foreign tissue. The invention is also useful for treating autoimmune diseases such as pemphigus vulgaris, rheumatoid arthritis, multiple sclerosis and systemilupus erythematosus. The present sequence is the DR2-IgG fusion protein encoding

G; 301 T; 0 U; 0 Other; Sequence 1446 BP; 414 A; 375 C; 356

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LysGluAsnAlaGlnLeuGluTrpGluLeuGlnAlaLeuGluLysGluLeu---AlaGln 259
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                   ProAsnLeuLeuGlyGlyProSerValPheIlePheProProLysIleLysAspValLeu
                                                                                                                       CCTAACCTCTTGGGTGGACCATCCGTCTTCATCTTCCCTCCAAAGATCAAGGATGTACTC
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                                                                  GCAGCATCTGAGCCCCAGAGGCCCACAATCAAGCCCTGTCCTCCATGCAAATGCCCAGCA
                                                                                                                                                                                                               AspValGlnIleSerTrpPheValAsnAsnValGluValHisThrAlaGlnThrGlnThr
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                                                      AlaAlaSerGluProArgGlyProThrileLysProCysProCysLysCysProAla
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                                                                                                                                                                                                   Multimeric complex for treating autoimmune diseases, comprises first and second single chain MHC class II molecules, each comprising alphal and betal domain linked through amino acid linker and multimerization domain.
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459 ArgvalGluLysLysAsnTrpValGluArgAsnSerTyrSerCysSerValValHisGlu
                                                                                                            Other,
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23-JAN-2001; 2001US-0264003P.
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P-PSDB; ABBS6463.
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1884 ACAGTCCCAGAAGTA-----TCATCTGTCTTCATCTTCCCCCAAAGCCCAAGGAT 1734

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rcent Similarity: st Local Similarity:

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298 ValLeuMetIleSerLeuSerProIleValThrCyBValValValAspValSerGluAsp 317

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                                                                                                                                        The invention relates to a multimeric complex comprising a first recombinant single chain major histocompatibility complex (MHC) class II molecule and a second recombinant single chain MHC class II molecule, each comprising an alphal domain and a betal domain linked through an amino acid linker and a multimerisation domain. The first and the second molecule are linked through the multimerisation domain to form a multimeric complex, The complex is useful for treating autoimmune diseases. It is useful for treating insulin dependent diabetes, multiple sclerosis, myasthenia gravis, pernicious anaemia, autoimmune encephalomyellitis (EAE), rheumatoid arthritis and systemic lupus erythematosus. The present sequence encodes a single chain MHC class II molecule of the invention
                                              Multimeric complex for treating autoimmune diseases, comprises first and second single chain MHC class II molecules, each comprising alphal and betal domain linked through amino acid linker and multimerization domain.
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WPI; 2001-616371/71.
P-PSDB; ABB56459.
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15-MAY-2000; 2000US-0204249P.
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Multimeric complex for treating autoimmune diseases, comprises first and second single chain MHC class II molecules, each comprising alphal and betal domain linked through amino acid linker and multimerization domain.

Disclosure; Page 95; 147pp; English.

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WPI; 2001-616371/71. P-PSDB; ABB56462.

22-MAR-2000; 2000US-0191274P. 15-MAY-2000; 2000US-0204249P. 23-JAN-2001; 2001US-0264003P.

(CORI-) CORIXA CORP

22-MAR-2001; 2001WO-US009616

The invention relates to a multimeric complex comprising a first recombinant single chain major histocompatibility complex (MHC) class II molecule and a second recombinant single chain MHC class II molecule, each comprising an alphal domain and a betal domain linked through an amino acid linker and multimerisation domain. The first and the second molecule are linked through the multimerisation domain. The first and the second multimeric complex, The complex is useful for treating autoimmune diseases. It is useful for treating insulin dependent diabetes, multiple sclerosis, myasthenia gravis, pernicious anaemia, autoimmune erocephalomyelitis (EAB), rheumatoid arthritis and systemic lupus erythematosus. The present sequence encodes a single chain MHC class II molecule of the invention

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Mouse; MHC; major histocompatibility complex; MHC class II; multimer; single chain; immunosuppressive; antidiabetic; antilnflammatory; antiannemic; antirheumatory; antiarthritic; neuroprofective; vaccine; autoimmune disease; insulin dependent diabetes; multiple sclerosis; myasthenia gravis; pernicious anaemia; autoimmune encephalomyelitis; rheumatoid arthritis; systemic lupus erythematosus; ss.
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ABI99032 standard; cDNA; 2059
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ımmunogenic compositions comprising Flt-3 ligand encoding polynucleotide
and one or more antigen, or cytokine encoding polynucleotides, useful for
suppressing tumor growth and for treating autoimmune diseases (e.g.
rheumatoid arthritis).
                                                                                                                                                                                                                                                                                                                                                                                                                                  The present sequence is that of patient-specific bicistronic chimeric idiotype VR1642 (plasmid VAXID), which is used to treat B-cell lymphoma patients. The plasmid includes the cytomegalovitus immediate-early promoter, enhancer and 5' untranslated sequences, driving the expression of mouse-human chimeric immunoglobulin light and heavy chain sequences. The human light and heavy chain variable regions are derived from B-cell lymphoma cell line RAMOS. The transcriptional terminator region includes polyA and termination signals from the bovine growth hormone gene. According to the invention, co-administration of VR1642 with a plasmid see AAR30314) encoding human Fme like tyrosine kinase (Flt-3 ligand) provides a means of treating a patient with B-cell lymphoma. (Updated on 11-SEP-2003 to standardise OS field)
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       GlnAsnIleAlaAlaGluLysHisAsnLeuGlyIleLeuThrLysArgSerAsnPheThr
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The present sequence is that of the cloning cassette of a tri-cistronic mAb17-1A expression and dihydrofolate reductase selection construct for use in the production of recombinant 1962a mAb17-1A antibody. In examples from the invention, comparison of the original, hybridoma-derived immunisation antigen 17-1A and recombinantly expressed mAb17-1A from CHO cimmunisation antigen not immunological differences in rhesus monkey critals. Both formulations showed identical kinetics building up the immunisation antigen and target antigen specific immune response, and raised 190 and 19M titres were similar. Recombinant mAb17-1A is an example of an immunogenic recombinant antibody of the invention designed for immunisation of primates and comprising at least a part of a murine 192a subtype amino acid sequence with mammalian glycosylation.

Recombinant DNA methods are used to produce the immunogenic antibody in a standardised manner. The humoral immune response induced by the 1962a immunogenic antibodies of the invention is significantly improved in terms of the quantity of specific antibody induced by the patients and the specificity against selected targets and epitopes. The improved
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immune response is dependent on the glycosylation pattern of the antibody. Recombinant antibody expressed in hamster or human cells were shown to have a similar immunogenicity as antibody expressed by murine hybridoma cells. This is of particular relevance for antibodies that are to be used for immunisation purposes. The antibody may have a murine amino acid sequence or any other mammalian amino acid sequence that is combined with the murine igg2a part. Preferably mammalian sequences are human, humanized, human/murine chimeric or murine sequences. The antibody may also be an anti-idiotypic antibody or a mimotopic Abl antibody. The igg2a immunogenic recombinant antibody can be directed against a tumour associated antigen. The invention also provides vaccines comprising the igg2a immunogenic antibody. The vaccines may be used for the prophylaxis and therapy of cancer associated diseases, e.g. metastatic disease in cancer patients. The vaccine specifically modulates antigen-presenting cells in vivo or ex vivo, thus generating an immune response to the epitope that is targeted by the IgG2a immunogenic antibody. The preferred method of producing the antibody-expression construct containing at least an uncleotide sequence encoding a kappa light chain and antibody. The preferred sequence encoding a sappa light chain and a least a part of a murine IgG2a subtype amino acid sequence encoding at least a part of a murine IgG2a antibody. The kappa light chain and gamma heavy control of a single CMV promoter to produce an inteact antibody. The kappa light chain and gamma heavy control of a single chain are expressed in about equined aquantity, and antibody control of a new and antibody control of a new and antibody control of the control of a single chain and antibody control of a new and antibody control of a new and antibody control of the control of a single chain and samma heavy chain. concentrations of 5-300 ug/ml are achieved

Sequence 3973 BP; 1052 A; 1038 C; 993 G; 888 T; 0 U; 2 Other;

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3973 281 26 71 95	aaspglyaspglul      caaggcaaggcaa	ogluphe	1PheGluProGlnG ::: TACTGGGGCCAAG	ThrLysArgSerA	oLysSerProval -   :::      \CTGGCCCTGTGT	AsnilePheProP       GGTTATTTCCCTG	oglyvalTyrgluT        ::: rggrgrgcacaccT	rLeuThrPhelleP    :::  CTCAGT	euGluGluProVal
Length: Matches: Conservative: Mismatches: Indels: Gaps:	42 SerProGlyAspileGlyGlnTyrThrHisGluPheAspGlyAspGluLeuPheTyr 60   1::	61 ValAspLeuAspLysLysLysThrValTrpArgLeuProGluPhe	76	88 GlnAsnIleAlaAlaGluLysHisAsnLeuGlyIleLeuThrLysArgSerAsnPheThr 	108 ProAlaThrAenGluAlaProGlnAlaThrValPheProLysSerProVal 124	125 LeuLeuGlyGlnProAsnThrLeuIleCysPheValAspAsnIlePheProProValIle 144	145 ABNIJEThrTrpLeuArgABRASerValThrAspGlyValTyrGluThrSerPhe 164   1873 ACCTTGACCTGGAACTCTGGATCCTGTCCAGTGGTGTGCACCTCCCAGCT 1926	165 LeuValAsnargAspHisSerPheHisLysLeuSerTyrLeuThrPheIleProSerAsp :::::	185 AspAspIleTyrAspCysLysValGluHis-TrpGlyLeuGluGluProValLeuLysHi         :::
finment Scores: 6.48e-104 Lengt d. No.: 1328.50 Match 10: 1328.50 Conse cent Similarity: 65.0\$ Conse t Local Similarity: 59.5\$ Misma 1:Y Match: 13.04.05 Match: 1.166.2 Match: Match: 1.166.2 Match: M	YABDIleGlyGli	uAspLysLysLysTh: ATCCTCCAGCACTGC	TTTCTGTGCAAGAGA	eAlaAlaGluLysHi ::::    CTCTGCAGCCAAA	rAsnGlualaProGl:        -ACAACAGCCCCA	yglnProAsnThrLe   	rTrpLeuArgAsnSe        AACTC	nArgAspHisSerPh      GTCTGACCT	eTyrAspCysLysVa
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                                                             rThrThrAlaProSerAlaGlnLeuGluLysGluLeuGlnAlaLeuGluLysGluAsnAl
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Query Match:

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New immunoglobulin hybrid proteins - with immunoglobulin fragments linked to dimeric protein, for diagnostic or therapeutic use.
anti-snake small neurotoxin antibody; heavy chain; 19G2; immunoglobulin; bispecific bivalent antibody; cell-targetting; cytotoxic agent; ss.
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Example 1; Fig 3A; 37pp; French.

A fragment of the heavy chain (VH + CHI) from the anti-snake small neurotoxin monoclonal antibody M(alpha)2-3 was PCR-amplified from hybridoma-derived CDNA using primers AAQ48039 and AAQ48040. A light chain fragment (VL + CL) was amplified from the same source using primers AAQ48041 and AAQ48042. The two amplified fragments were inserted into the same vector; the H-chain fragment was inserted (in-frame) between codons 6-7 of the photh contained a ploth S-D sequence, a signal peptide and the first 6 codons of photh. The cassette was positioned between the termination codon and the transcription termination sequence of photh. The fusion construct is expected to to encode a hybrid protein comprising two identical Ab-derived units. The invention also covers hybrid proteins containing two different Ab-derived units (1.e. to produce bispecific antibodies). When a toxic protein is used in place of photh, the hybrid molecules can be used as cell-targetting therapeutic agents. (Updated on 10-MAR-2003 to add missing of field.) (Updated on 25-MAR-2003 to correct PN field.)

Sequence 1581 BP; 435 A; 448 C; 373 G; 325 T; 0 U; 0 Other; ignment Scores:

ad. No.:
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1324 50
Antches:
1286
Conservative: 31
St Local Similarity: 59.0\$
Mismatches: 81

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ò	>	34	PheTyrGlyThr	ThrValTyrGln	SerProGlyAsp	11eGly	4.7
g	۵	211		TGGTGAAGCAG	AAGCCTGGACAGGG	ACTTAAATGGATTGGATG	270
ò	>-	8		Gln	TyrThrHisGluPh	GlnTyrThrHisGluPheAspGlyAspGluLeuPheTyr	09
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ò	>-	61		LysLysLysThr	ValTrpArgLeu	ValAspieuAspiysiysiysThrValTrpArgleuProGluPhe	75
g	۵	331	GTAGACACATCC	TCCAGCACAGCC	racargcagcicag	CAGCCTGACATCTGAGGACACT	390
ò	>-	76		LeuPheGluPro	GlnGlyGlyLeuGl	GlyGlnLeulleLeuPheGluProGlnGlyGlyLeuGlnAsnIleAlaAlaGluLysHis	95
qq	۵	391	GCTGT-CTATTT	CTGTGCAAGAGC	TATGGGGGCTAC	GGCTAC	431
È	>-	96		LeuThrLysArg	-SerAsnPheThrP	AsnLeuGlylleLeuThrLysArg-SerAsnPheThrProAlaThrAsnGluAlaProGl	115
g	۵	432		creecccaae	caccacrcrcacad	rctcctcagccaaacaacagc	491
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ò	>-	152	rLysSerValTh	rAspGlyValTy	rGluThrSerPheL	euValAsnArgAspHisSerPh	172
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ò	>	172	eHisLysLeuSe	rTyrLeuThrPh	elleProSerAspA	spAsplleTyrAspCysLysVa	192
q	۵	99	CTACACCCTCAG	cag-crcagr		GACTGTAACCT	691
ò	>-	192	lGluHis-TrpG	lyLeuGluGluP	GluHis-TrpGlyLeuGluGluProValLeuLysHis  	192 1GluHis-TrpGlyLeuGluGluProValLeuLysHisTrpGluProGluIleProAlaP 2 	ᅼ
g	۵	692	caaccaccree-	1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1		- O	0
ò	>-	212	roMetSerGlub	euThrGluThrG	lyGlyGlyGlySer'	roMetSerGlubeuThrGluThrGlyGlyGlyGlySerThrThrAlaProSerAlaGlnL	232
g	۵	704		TCACC	TGCAATGTGGCC	caccedeaagcadeaccaagg	754
ò	>	232		euGlnAlaLeuG	luLysGluAsnAla	3lnLeuGluTrpGluLeuGlnA	252
a	۵	755		TT			168
ò	>-	252		luLeuAlaGlnAle	laAlaSerGluPro	erGluProArgGlyProThr1leLy8ProC	272
đ	۵	769			GAGCCC	AGAGGCCCACAATCAAGCCCT	196
ò	>-	272		ysCysProAlaP	roAsnLeuLeuGly	ysproproCysLysCysProAlabroAsnLeuLeuGlyGlyProSerValPhellePheP	292
g	۵	797		AATGCCCAGCAC	craaccrcrrdger	saaccarccarcrrcarcrrcc	958
ò	>-	292		ysAspValLeuM	etlleSerLeuSer	roProLysileLysAspValLeuMetileSerLeuSerProlleValThrCysValValV	312
음	۵	857	_	AGGATGTACTCA	rearcrecteage	ccaragreacarereres	916
8 1	٠.	312		luaspaspProa	SpvalGlnIleSer	alagpvalSerGluAgpAgpProAgpValGlnIleSerTrpPheValAgnAsnValGluV	332
â	۵	917		AGGATGACCCAG	AIGICCAGAICAGC	IGGITIGICAACAACGIGGAAG	9 1
È	>-	332		InThrGlnThrH	18ArgGluAspTyr	alHisThralaGlnThrGlnThrHisArgGluAspTyrAsnSerThrLeuArgValValS	352
셤	ڡ	977		AGACACAAACCC	ATAGAGAGGATTAC	AACAGTACTCTCCGGGTGGTCA	1036

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                                                                                                                                          ACAACAAAGACCTGCCAGCGCCCATCGAGAGCCATCTCAAAACCCAAAGGGTCAGTAA 1156
                                                                                                                                                                                                                                     GAGCTCCACAGGTATATGTCTTGCCTCCACAGAAGAAGATGACTAAGAAACAGGTCA 1216
                                                                                                                                                                                                                                                                                                               CTCTGACCTGCATGGTCACAGACTTCATGCCTGAAGACATTTACGTGGAGTGGACCAACA 1276
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prion disease; 44BlH; ds; therapeutic; monoclonal antibody; heavy chain.
                                                                                                                                                                                                                                                                                                                                                                                                          ACGGGAAAACAGAGCTAAACTACAAGAACACTGAACCAGTCCTGGACTCTGATGGTTCTT
                                                                                                                                                                                                                                                                                                                                                                                                                                                                     yrPheMetTyrSerLysLeuArgValGluLysLysAsnTrpValGluArgAsnSerTyrS
erAlaLeuProIleGlnHisGlnHsGlnAspTrpMetSerGlyLysGluPheLysCysLysValA
                                                                                             snAsnLysAspLeuProAlaProIleGluArgThrIleSerLysProLysGlySerValA
                                                                                                                                                                                      rgAlaProGlnValTyrValLeuProProProGluGluGluGluMetThrLysLysGlnValT
                                                                                                                                                                                                                                                                                      hrLeuThrCysMetValThrAspPheMetProGluAspIleTyrValGluTrpThrAsnA
                                                                                                                                                                                                                                                                                                                                                                                  snGlyLysThrGluLeuAsnTyrLysAsnThrGluProValLeuAspSerAspGlySerT
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       ACTICATGIACAGCAAGCIGAGAGTGGAAAAGAAGAACTGGGTGGAAAGAAATAGCTACT
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      erCysSerValValHisGluGlyLeuHisAsnHisHisThrThrLysSerPheSerArgT
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cc as that of the above nucleotide sequence, a nucleotide sequence, which is a mutant of the above sequences or a nucleotide sequence that is complementary to the above sequences and that hybridizes under stringent conditions with the above sequences and that hybridizes under stringent conditions with the above sequence; and an antibody light chain gene having SEQ ID No: 2, 4, 6, 31, 33 and 35, a nucleotide sequence consisting of degenerate genetic code, which encodes a polypeptide same as that of the above nucleotide sequence, an uncleotide sequence, which is a mutant of the above sequences, or a nucleotide sequence, which is a mutant of the above sequences, or a nucleotide sequence, which is conditions with the above sequences and that hybridizes under stringent conditions with the above sequences and that hybridizes under stringent conditions with the above sequences and that hybridizes under stringent conditions with the above sequences and that hybridizes under stringent conditions with the above sequences and that hybridizes under stringent conditions and constant region of antibody of animal other than of another than constant prion proliferation inhibition activity, comprising transducing a gene that provides abnormal prion proliferation inhibition activity to continized for treating prion disease; use of a mesenchymal cill for cutilized for treating prion disease; use of a mesenchymal cell for treating prion disease; use of a mesenchymal cell for preparing a conducing an agent for delivering a substance to the lesioned region of prion disease; comprising utilizing mesenchymal cell. (I) is useful for treating prion disease. (II) (IV) or (IV) is useful for treating prion disease or delivering a substance to the lesioned region of prion disease. (III) (IV) or (IV) is useful for treating prion disease. (IV) is useful for treating prion disease. (IV) is useful for treating prion disease. (IV) is useful for treating prion disease. (IV) is useful for treating prion disease. (IV) is useful for preparing a chic 165 185 126 LeuGlyGlnProAsnThrLeuIleCysPheValAspAsnIlePheProProValIleAsn 145 673 727 205 784 AlaThrAsnGluAlaProGlnAlaThrValPheProLysSerProVal------Leu 125 614 ACTGGCTCCTCGGTGACTCTAGGATGCCTGGTCAAGGGTTATTTCCCTGAGCCAGTGACC 166 ValAsnArgAspHisSerPheHisLysLeuSerTyrLeuThrPheIleProSerAspAsp 146 IleThrTrpLeuArgAsnSerLysSerValThrAspGlyValTyrGluThrSerPheLeu CTGCAGTCTGAC-----CTCTACACCCTCAGCAG-CTCAGT--------186 AspileTyrAspCysLysValGluHis-TrpGlyLeuGluGluProValLeuLysHisTr directly from WIPO at ftp.wipo.int/pub/published\_pct\_squences Sequence 1560 BP; 413 A; 424 C; 388 G; 335 T; 0 U; 0 Other; 1560 269 19 39 65 Length: Matches: Conservative: Mismatches: Indels: Gaps: US-10-048-116B-2 (1-495) x AED19725 (1-1560) 7.55e-104 1321.50 73.7% 68.8% Percent Similarity: Best Local Similarity: Alignment Scores: 109 260 674 728 763 Query Match: DB: ŝ 셤 ઠે g ò 셤 ò 셤 à 셤

816

-------TGCAGTCCATCACC-----TGCAATGTGGCCCA

pGluProGluIleProAlaProMetSerGluLeuThrGluThrGlyGlyGlyGlySerTh

205

8 8 8 8 8

rThralaProSerAlaGlnLeuGluLysGluLeuGlnAlaLeuGluLysGluAsnAlaGl

cccecchagcaccaaggrggachagaaarr---

The invention describes an agent (I) for treating prion disease or delivering a substance to the lesioned region of prion disease, comprising a meeenchymal cell. Also described are: a nucleic acid (II) having an anti-prion antibody gene comprising: an antibody heavy chain gene having SEQ ID No: 1, 3, 5, 30, 32 and 34, a nucleotide sequence consisting of a degenerate genetic code, which encodes a polypeptide same

Claim 4; SEQ ID NO 5; 34pp; Japanese.

nLeuGluTrpGluLeuGlnAlaLeuGluLysGluLeuAlaGlnAlaAlaSerGluProAr 265

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----GAGCCCAG 858
                                                                                                                                                    285 yProSerValPheIlePheProProLysIleLysAspValLeuMetIleSerLeuSerPr 305
                                                                                                                                                                                  ACCATCCGTCTTCATCTTCCTCCTCCAAAGATCAAGGATGTACTCATGATCTCCTGAGCCC
                                                                                                859 AGGGCCACAATCAAGCCCTGTCCTCCATGCAAATGCCCAGCACCTAACCTCTTGGGTGG
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                                              gGlyProThrileLysProCysProProCysLysCysProAlaProAsnLeuLeuGlyGl
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antiparkinsonian; anticonvulsant; protein production; therapeutic; pharmaceutical; amyloidosis; metabolic disorder; cerebrovascular ischemia; cardiovascular disease; neurological disease; brain injury; injury; spinal cord injury; vulnerary; dementia; genetic disorder; Creutzfeldt Jakob disease; Huntingtons chorea; motor neurone disease; infection; antiinflammatory; inflammatory; inflammatory; antiinflammatory; antiinflammatory; antiinflammatory; antiinflammatory; antiinflammatory; antiinflammatory; antibody; heavy chain; ds. neuroprotective; nootropic; cerebroprotective; vasotropic; Anti-Nogo-antibody 2A10 heavy chain polynucleotide. ВЪ AEB21727 standard; DNA; 1407 (first entry) 08-SEP-2005 AEB21727;

The invention describes a method of modulating production of an amyloidogenic peptide comprising contacting a cell which is expressing contacting a cell which is expressing contacting accell which is expressing contacting in the product antagonist. Also described are: use of a Nogo antagonist. Also described are: use of a Nogo antagonist in the manufacture of a medicament for the treatment or prophylaxis of a disease involving amyloidosis; and a method of treatment or prophylaxis of Alzheimer's disease comprising administering to the human in need an anti-Nogo antibody. The invention is used for modulating production of amyloidogenic peptide in, e.g. Alzheimer's disease, stroke, traumatic brain injury and spinal cord injury, fronto-temporal dementias, peripheral neuropathy, Parkinson's disease, Huntington's disease, creutzfeldt-Jakob disease, amyotropic lateral sclerosis, multiple sclerosis, or inclusion body myositis. The invention provides an unexpected route for therapeutic intervention in particularly Alzheimer's disease, this sequence represents an anti-Nogo-antibody 2AlO heavy chain Modulating production of amyloidogenic peptide in, e.g. Alzheimer's disease, by contacting cell and Nogo polypeptide with Nogo antagonist Sequence 1407 BP; 378 A; 396 C; 346 G; 287 T; 0 U; 0 Other; Example 4; SEQ ID NO 49; 53pp; English 22-DEC-2003; 2003GB-00029684. 22-DEC-2003; 2003GB-00029711. 20-DEC-2004; 2004WO-GB005343 (GLAX ) GLAXO GROUP LTD Prinjha RK; WPI; 2005-522181/53 polynucleotide 07-JUL-2005. Hussain I, 

Alignment Scores:

ò 셤 ò g 8 a 8

Length: Matches:

ProGlnAlaThrvalPheProLysSerProVal------LeuLeuGlyGlnProAsn 130 237 357 ------ 423 93 Grigaagcagaccigaacaagcccrrgagragarricgaararraarccragcaarggr 48 -----GlnTyrThrHisGluPheAspGlyAspGluLeuPheTyrValAspLeuAspLys AGCACAGCCTACATGCAGCTCAGCCTGACATCTGAGGACTCTGCGGTCTATTATTGT LysHisAsnLeuGlyIleLeuThrLysArgSerAsnPheThrProAlaThrAsnGluAla GlupheGlyGlnLeuIleLeuPheGluProGlnGlyGlyLeuGlnAsnIleAlaAlaGlu LysLysThrValTrpArgLeuPro------ValTyrGlnSerProGlyAsp------ileGly-----1407 285 26 68 108 Conservative: Mismatches: Indels: AAA----Gaps: (1-1407)US-10-048-116B-2 (1-495) x AEB21727 8.87e-104 1320.00 64.0% 58.6% Percent Similarity: Best Local Similarity: 39 238 99 74 358 94 412 Query Match: DB:

|||| :::|||:::||| CCA-----TCGGTCTATACAACTACAACTGGCTCCTCGGTG 477

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6. B 쉱 ò

Mus sp.

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                                                                                                                                                            AsnSerLysSerValThrAspGlyValTyrGluThrSerPheLeuValAsnArgAspHis 170
                                                                                                                                                                                                                SerPheHisLysLeuSerTyrLeuThrPheIleProSerAspAspAspIleTyrAspCys 190
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ThrLeuileCysPheValAspAsnilePheProProValileAsnileThrTrpLeuArg
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New antibody or its functional fragment that binds with and neutralizes human neurite outgrowth useful for treating or prophylaxis of stroke and other neurological disease e.g. traumatic brain injury, spinal cord
                                                                                                                                                                                                                         disease
                                                                                                                                            cerebroprotective, vasotropic, neuroprotective, vulnerary, nootropic antiparkinsonian; anticonvulsant; neuroleptic; antibody engineeting; pharmaceutical; cerebrovascular ischemia; cardiovascular disease; neurological disease; brain injury; injury; spinal cord injury; Alzheimers disease; degeneration; dementia; neuropathy; parkinsons disease; Huntingtons chorea; genetic disorder; multiple sclerosis; immune disorder; cetuzzfeldt Jakob disease; infection; schizophrenia; psychiatric disorder; motor neurone disease; cns-gen.; muscular-gen.; ds.
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                                                                                                                        antibody 2A10 heavy chain polynucleotide SEQ ID NO 49.
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Wilson PA;
                                                        BP
CCGGACTCCGGGTAAA 1398
                                                       DNA; 1407
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22-DEC-2003; 2003GB-00029711
                                                                                                    (first entry)
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Plumpton C, Prinjha RK,
                                                                                                                                                                                                                                                                                                                                                                                     (GLAX ) GLAXO GROUP LTD.
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                                                         AEB08761 standard;
                                                                                                                                                                                                                                                                                   WO2005061544-A2
                                                                                                    08-SEP-2005
                                                                                                                                                                                                                                                              Unidentified
            1383
                                    RESULT
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The invention describes an antibody (A1) or its functional fragment, that bids with and neutralizes human neurite outgrowth (NOGO). Also described are: providing a second vector encoding a heavy chain of the antibody; corporating a mammalian host cell with the first and second vectors: transfecting a mammalian host cell with the first and second vectors: conditions permissive to the secretion of the antibody from the host cell into the culture media; recovering (and optionally purifying) the into the culture media; recovering axonal sprouting involving contacting a secreted antibody; and promoting axonal sprouting involving contacting a human axon with an anti-NOGO antibody. The antibody is useful in the preparation of a medicament for treating or prophylaxis of stroke and cother neurological disease, disorders (e.g. traumatic brain injury, spinal cord injury, Alzheimer's disease, frontotemporal dementias (tauopathies), cord injury, Alzheimer's disease, frontotemporal dementias (tauopathies), cord injury, parkingon's disease (CDD), Schizophrenia, amyotrophic lateral sclerosis (ALS), inclusion body myositis. The amyotrophic lateral sclerosis (ALS), inclusion body myositis. The neurological diseases/disorder: This sequence represents an antibody 2AlO neurological diseases/disorder: This sequence represents an antibody 2Alo are the creation of recombinant anti-NOGO

Example 3; SEQ ID NO 49; 143pp; English.

injury, Alzheimer's disease.

Sequence 1407 BP; 378 A; 396 C; 346 G; 287 T; 0 U; 0 Other;

antibodies.

Aliqnment Scores:

rArgThrProGlyLys 495

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423
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AACTCTGGATCCCTGTCCAGTGTGCACACCTTCCCAGCTGTCCTGCAGTCTGAC--- 588
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 1407
285
26
68
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13
 Length:
Matches:
Conservative:
Mismatches:
Indels:
Gaps:
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1262 1022 430 ySerTyrPheMetTyrSerLysLeuargValGluLysLysAsnTrpValGluArgAsnSe 470 370 962 CAACAACGGGAAAACAGGGTAAAACTACAAGAACACGGAACCAGGCCTGGACTCTGATGG 963 GGTCAGTGCCCTCCCCATCCAGCACCAGGACTGGATGAGTGGCAAGGAGTTCAAATGCAA |ValSerAlaLeuProlleGlnHisGlnAspTrpMetSerGlyLysGluPheLysCysLy rAsnAsnGlyLysThrGluLeuAsnTyrLysAsnThrGluProValLeuAspSerAspGl 1GluvalHisThrAlaGlnThrGlnThrHisArgGluAspTyrAsnSerThrLeuArgVa GGAAGTACACACACTCAGACACAAACCCATAGAGAGGATTACAACAGTACTCTCCGGGT ccedacrccederaa 1398 rArgThrProGlyLys 495 350 1143 1203 450 1383 330 903 410 430 490 8 è 셤 ያ ያ 8686 ઠે В ઠે 셤 ઠે g ઠે

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Recombinant protein; alpha chain; beta chain; WHC; immunoglobulin; major histocompatibility complex; Rc region; antigen; T lymphocyte; immunostimulant; vaccine; infection; tumour; ss.
                                                                                                                                                                                                                                                                                                                                                       DNA encoding a fusion protein comprising an alpha chain of MHC.
                 ADX26089
AAT06285
AAQS6919
AAT17587
AAT86988
                                                                                                          AAT60704
AAT17580
AAT17580
AAT17580
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AAT86987
ACA60742
AB199022
AB199023
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AB199031
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ADV43942
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AEB23393
AAI58351
                                                                                                                                                                                                                                                                                                                                                                                                         Location/Qualifiers
1. .1482
                                                                                                                                                                                                                                                                                                                AAF55098 standard; DNA; 1484 BP
                                                                                                                                                                                                                                                                                                                                          15-MAY-2001 (first entry)
                                    WO200109194-A1
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May 31, 2006, 22:51:28 ; Search time 557.439 Seconds (without alignments) 5215.692 Million cell updates/sec
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                                                                  US-10-048-116B-2_COPY_1_278
1496
1 MPCSRALILGVLALNTMLSL......QAASEPRGPTIKPCPPCKCP 278
                                                                                                                                                                                                                                                                                                                                                                                                                                                Description
      GenCore version 5.1.8
Copyright (c) 1993 - 2006 Biocceleration Ltd
                             nucleic search, using frame_plus_p2n model
                                                                                                                                                al number of hits satisfying chosen parameters:
                                                                                                                                    5244920 segs, 3486124231 residues
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Maximum Match 100%
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Ygapop 10.0, Ygapext
Fgapop 6.0, Fgapext
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28-JUL-2000; 2000WO-FR002193

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5 AAF55098 12 ADQ31223 2 AAV12067

1484 777 4713

100.0 81.2 79.0

1496 1214.5 1182

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99FR-00009862

29-JUL-1999;

Aca60743 Mouse MRC
Aca6098 Alphalalp
Aba30054 IAB alpha
Abi30054 Murine bC
Aca60700 CDNA enco
Aca60704 Murine bC
Aca60744 Mouse MRC
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(CNRS ) CNRS CENT NAT RECH SCI.
                                           Glaichenhaus N, Malherbe L;
                                                                                      WPI; 2001-182944/18.
P-PSDB; AAB67480.
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t C New soluble recombinant protein, useful e.g. as immunostimulant, comprises dimeric major histocompatibility complex molecule fused immunoglobulin Fc region.

Example 1; Page 31-33; 43pp; French

The specification describes soluble recombinant proteins that comprise at least a dimer formed from the alpha and beta-chains of MHC (major histocompatibility complex) Class I and II molecules in which at least one chain has, attached to its C-terminus at least part of the Fc region of an immunoglobulin. The recombinant proteins, when linked to an antigenic peptide, are used to count and/or purify antigen-reactive T lymphocytes and to characterize their phenotype, e.g. in preclinical evaluation of vaccines. They are also used as immunostimulants, particularly for vaccine development (against infections and tumours), to count and determine phenotype of autoreactive T cells in subjects with, or at risk of developing, autoimmune diseases, e.g. for staging or evaluating treatments, and (to purify and/or enrich Ag-reactive T cells from cell cultures or patient samples, for use in subsequent curative or preventative cellular therapy. The present sequence encodes a recombinant protein of the invention, comprising an alpha chain of MHC molecules

Sequence 1484 BP; 414 A; 394 C; 362 G; 314 T; 0 U; 0 Other;

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21		sluaspaspileglu 	nlaaspHisvalGly                 gccgaccacgraggc	CysglyGlyGluAspAspIleGluAlaAspHisValGlyPheTyrGlyThrThrValTyr 40 
41	_	31yAspileGlyGlr 	1TyrThrHisGluPhe 	GlnserProglyAspileGlyGlnTyrThrHisGluPheAspGlyAspGluLeuPheTyr 60 
181		ABDLYBLYBLYBThz 	.ValTrpArgLeuPro 	ValaspLeuaspLysLysLysThrValTrpArgLeuProGlubheGlyGlnLeuIleLeu 80 
81		slnglyglybeuglr 	nasnilealaalaglu                 naacatagcrgcagaa	PhegluProglnGlyGlyLeuglnAanIleAlaalaGluLy8HisAsnLeuglyIleLeu 100 
101		SerAsnPheThrPrc 	oalathrasnGluala                 \GCTACCAATGAGGCT	ThriygargseraenPheThrProalaThrAenGlualaProglnalaThrValPhePro 120 
121		ValleuleuGlyGlr                   5rGCrGCrGGGrCAC	nProdenThrLeuIleC) 	LysserProvalLeuLeuGlyGlnProAsnThrLeuIleCysPheValAspAsnIlePhe 140 
141			oleuArgAsnSerLys 	Proprovalileaenilethrirpleuargaenseriyeservalthraepglyvaltyr 160 

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Peptide-Class II major histocompatibility complex (MHC) composite, useful for detecting antigen specific CD4+ T-cell, comprises antigen peptide containing epitope of mucous membrane invasive protein, and extracellular region of MHC.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         The invention relates to a novel class II major histocompatibility complex (MHC) antigenic peptide composite comprising a peptide containing the T-cell antigenic determinant of a mucous membrane invasive protein and the extracellular region of a class II MHC molecule or at least part of the extracellular region of the class II MHC molecule having an amino acid sequence comprising one or more deletions, substitutions or
                                                         200
                                                                                                                     220
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                                                                                                                                                                                                 260
                                                                                                                                                                                                                                                                   721 AAGGAAAATGCACGAGCTCGAATGGGAGTTGCAAGGAAATGGAAAAGGAACTGGCTCAGGCA 780
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          class II major histocompatibility complex; MHC; CD4+ T-cell detection; flow cytometry; mucous membrane invasive antigen; I-Ab(alpha)-leucine zipper fusion; 88; gene.
GluThrSerPheLeuValAsnArgAspHisSerPheHisLysLeuSerTyrLeuThrPhe
                                                                                                                                                                             GlyGlyGlySerThrThrAlaProSerAlaGlnLeuGluLysGluLeuGlnAlaLeuGlu
                                                           11eProSerAspAspAspIleTyrAspCysLysValGluHisTrpGlyLeuGluGluPro
                                                                          GTTCTGAAACACTGGGAACCTGAGATTCCAGCCCCCATGTCAGAGCTGACAGAAACTGGA
               481 GAGACCAGCTTCCTCGTCAACCGTGACCATTCCTTCCACAAGCTGTCTTATCTCACCTTC
                                                                                                                     ValleuLysHisTrpGluProGluIleProAlaProMetSerGluLeuThrGluThrGly
                                                                                                                                                                                                                                       LysGluAsnAlaGlnLeuGluTrpGluLeuGlnAlaLeuGluLysGluLeuAlaGlnAla
                                                                                                                                                                                                                                                                                                AlaSerGluProArgGlyProThrIleLyBProCyBProCyBLyBCyBPro 278
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                /*tag= a
/product= "Class II MHC-related I-Ab(alpha)-leucine
zipper (LZ) fusion protein"
                                                                                                                                                                                                                                                                                                                     781 GCATCTGAGCCCAGAGGGCCCACAATCAAGCCCTGTCCTTCCATGCAAATGCCCA 834
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Class II MHC-related I-Ab(alpha)-leucine zipper (LZ) fusion cDNA.
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 (SENT-) SENTAN KAGAKU GLJUTSU INCUBATION CENT
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Location/Qualifiers
1. .777
                                                                                                                                                                                                                                                                                                                                                                                           ADQ31223 standard; cDNA; 777 BP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       03-DEC-2003; 2003JP-00404367.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     03-DEC-2002; 2002JP-00351818.
                                                                                                                                                                                                                                                                                                                                                                                                                                                      07-OCT-2004 (first entry)
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additions. The molecule of the invention may be useful for detecting an antigen-apecific CD4+ T-cell by flow cytometry and for presenting a microorganism-derived mucous membrane invasive protein as an antigen. The method of the invention enables efficient detection of antigen-specific activation of CD4+ T-cells in the amedians membrane. The current sequence is that of the class II major histocompatibility complex-related I-AD(alpha)-leucine zipper (LZ) fusion cDNA of the invention.

Sequence 777 BP; 193 A; 207 C; 195 G; 182 T; 0 U; 0 Other;

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| IleProSerAspAspAspIleTyrAspCysLysValGluHisTrpGlyLeuGluGluFro 200
                                                                                                                                                                                                                                                                                                                                       61 TGTGGAGGTGAAGACGACATGAGGCCGACCACGTAGGCACCTATGGTATAAGTGTATAT 120
                                                                                                                                                                                                                                                                                                                                                                                                                              121 CAGTCTCCTGGAGACATTGGCCAGTACACATTTGAATTTGATGGTGATGATGTTCTTTAT 180
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 ProProvallleAsnIleThrTrpLeuArgAsnSerLysSerValThrAspGlyValTyr 160
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        ValleuLysHisTrpGluProGluIleProAlaProMetSerGluLeuThrGluThrGly 220
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             GlyGlyGlySerThrThrAlaProSerAlaGlnLeuGluLysGluLeuGlnAlaLeuGlu 240
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                                                                                                                                                                                                                                                  ATGCGGGGGGAGGACTCTGATTCTGGGGGTCCTCGCCCTGACCACCATGCTCAGCCTC
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                                                                                                                                                                                                                                                                                                              21 CysGlyGlyGluAspAspIleGluAlaAspHisValGlyPheTyrGlyThrThrValTyr
                                                                                                                                                                                                                            Met ProCysSerArgAlaLeuIleLeuGlyValLeuAlaLeuAsnThrMetLeuSerLeu
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                               No.:
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This nucleotide sequence comprises a PCR product obtained by amplification of mouse splenocyte cDNA using primers (see AAV12663 and AAV12064) designed for the amplification of IAd alpha chain full-length cONA. IAd beta chain cDNA (see AAV12068) has been similarly obtained. The cONA. IAd beta chain cDNA (see AAV12068) has been similarly obtained. The cIA sequences were cloned into metallothionein promoter (see AAV12062) - driven vector pRmHa-3 prior to sequencing. Major histocompatibility complex (MHC) class II IAd heterodimers were expressed at the call complex (MHC) class II IAd heterodimers were expressed at the call complex (MHC) class II Add heterodimers were expressed at the call complex (MHC) class II Add heterodimers were expressed at the call complex complex cells that have been transfected to produce MHC antigen presenting colles with one or more accessory molecules. The matrices are used to activate naive CD4+ T calls and to shift the ongoing activation state into a preferred differentiated population of Thi or Th2 calls. Applications include the treatment of autoimmune disease, es diabetes, multiple sclerosis, autoimmune thyroiditis, systemic lupus erythematosus, myasthenia gravis, Crohn's disease and inflammatory bowel disease, or an
                                                                                                                                                                                                                                                                                                                                                 Major histocompatibility class II antigen; MHC class II; T cell; T lymphocyte; Th1; Th2; activation; CD4+; antigen presenting cell; APC; autoimmune disease; diabetes; multiple sclerosis; autoimmune thyroiditis; systemic lupus erythromatosus; myssthenia gravis; Crohn's disease; inflammatory bowel disease; allery; athma; contact sensitivity; immunotherapy; therapy; IAd alpha chain; mouse; ds; circular; cyclic.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Synthetic antigen presenting cell for activating CD4+ T cells - usefutreat autoimmune disease, e.g. diabetes, multiple sclerosis, Crohn's disease and inflammatory bowel disease, or allergy, e.g. asthma and contact sensitivity.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Sequence 4713 BP; 1211 A; 1166 C; 1152 G; 1184 T; 0 U; 0 Other;
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Example 2; Page 92-94; 141pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           SR, Wingvist O, Karlsson L,
                                          BP
                                          CDNA; 4713
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               97WO-US008697
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           96US-0018175P.
                                                                                                                                                                                                                                                                                 Murine IAd alpha chain cDNA.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           (SCRI ) SCRIPPS RES INST.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       WPI; 1998-041895/04
                                     AAV12067 standard;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Mus musculus.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               22-MAY-1997;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           23-MAY-1996;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    W09746256-A1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    11-DEC-1997.
                                                                                                                                                                                                      08-JUN-1998
                                                                                                                 AAV12067;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Webb
AAV12067

XX AAV1

XX AAV1

XX AAV1

XX AAV1

XX AAV1

XX AAV1

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XX AV1



S

x AAV12067 (1-4713)

US-10-048-116B-2\_COPY\_1\_278 (1-278)

4713 219 0 0 0

Length:
Matches:
Conservative:
Mismatches:
Indels:

1.15e-107 1182.00 100.0\$ 100.0\$

Percent Similarity: Best Local Similarity:

Query Match:

Alignment Scores:

.. 9

Score:

Sheppard PO;

Deshpande S,

Gross JA,

95US-0005964P.

27-OCT-1995;

(ZYMO ) ZYMOGENETICS INC (ANER-) ANERGEN INC. Kindsvogel W, Reich EP,

```
878
   938
   180
  200
  758
  818
   LysSerProValLeuLeuGlyGlnProAsnThrLeuIleCysPheValAspAsnIlePhe 140
  ProProValileAsnileThrTrpLeuArgAsnSerLysSerValThrAspGlyValTyr 160
   579 CAGTCTCCTGGAGACATTGGCCAGTACACATGAATTTGATGGTGATGAGTTGTTCTAT 638
  698
   PheGluProGlnGlyGlyLeuGlnAsnIleAlaAlaGluLy8HisAsnLeuGlyIleLeu 100
  ThriyeArgSerAsnPheThrProAlaThrAsnGluAlaProGlnAlaThrValPhePro 120
  80
  9
  1059 GTTCTGAAACACTGGGAACCTGAGATTCCAGCCCCCATGTCAGAGGCTGACAGAAACT 1115
  Soluble, fusion; major histocompatibility complex; MHC; heterodimer; complex; antigen; binding groove; tolerance; autoantigen; disease; insulin dependent; diabetes mellitus; IDDM; antagonist; T cell; anergy; presenting cell; ds.
  ValleuLysHisTrpGluProGluIleProAlaProMetSerGluLeuThrGluThr 219
                  ValAspieuAspiysiysiysThrValTrpArgleuProGluPheGlyGlnLeuIleLeu
   GluThrSerPheLeuValAsnArgAspHisSerPheHisLysLeuShrPhe
   | IleProSerAspAspAspIleTyrAspCysLysValGluHisTrpGlyLeuGluGluPro
CysGlyGlyGluAspAspIleGluAlaAspHisValGlyPheTyrGlyThrThrValTyr
  GlnSerProGlyAsp1leGlyGlnTyrThrHisGluPheAspGlyAspGluLeuPheTyr
  Greeactrecataagaagaaactercreeagectrecreagtriceccaarteatacte
  cDNA encoding soluble fused MHC heterodimer:peptide complex pLJ23
   Location/Qualifiers
1. .1344
/*tag= a
   AAT60705 standard; cDNA; 1344 BP
   95US-00482133.
95US-00482133.
95US-00483241.
   96WO-US010102
  (revised)
(first entry)
  07-JUN-1995;
07-JUN-1995;
07-JUN-1995;
   Key
mat_peptide
  WO9640944-A2
   07-JUN-1996;
  27-AUG-2003
12-SEP-1997
   19-DEC-1996
   Mus sp.
Synthetic.
   201
     21
  41
   61
  639
   669
   101
  759
   121
   819
  141
  879
   161
   939
  181
   666
  AAT60705;
  81
```

```
The present sequence encodes a novel soluble fused major
histocompatibility complex (MHC) heterodimer:peptide complex, comprising
that and 2nd MHC domains, linked by a 5-25 residue linker, an antigenic
peptide able to associate with a peptide binding groove of the MHC
molecule, linked in frame to the 2nd domain by a 5-25 residue linker and
a DNA encoding a 3rd MHC domain linked in frame to the DNA encoding the
an be used to induce immunological tolerance in adults susceptible to,
or suffering from an autoantigen related disease, e.g. insulin dependent
clabetes mellitus (IDDN), by antagonising the binding of particular T
clabetes mellitus (IDDN), by antagonising the binding of particular T
clab and antigen presenting cells, to induce anergy (immunological non-
responsiveness) in the targeted T cell. As the heterodimers and
corresponding antigen are permanently linked into a single chain,
obviating the requirement for complex heterodimer truncation or
formation, the complex eliminates inefficient and non-specific peptide
loading. (Updated on 27-AUG-2003 to correct OS field.)
  ProValileAsnileThrTrpLeuArgAsnSerLysSerValThrAspGlyValTyrGlu 161
   Novel fused major histocompatibility complex:antigenic peptide complex - useful to induce tolerance to an autoantigen-related disease e.g. insulin -dependent diabetes mellitus.
   507
   567
  GlubroGlnGlyGlyLeuGlnAsnIleAlaAlaGluLysHisAsnLeuGlyIleLeuThr 101
   LysargSerasnPheThrProAlaThrAsnGluAlaProGlnAlaThrValPheProLys 121
   687
  141
   747
   61
  81
  SerProValLeuLeuGlyGlnBroAsnThrLeuIleCyaPheValAspAsnIlePhePro
  688 TCCCCTGTGCGCTGGGTCAGCCCCAACACCCTTATCTGCTTTGTGGACAACATCTTCCCA
   SerbroGlyAspileGlyGlnTyrThrHisGluPheAspGlyAspGluLeuPheTyrVal
  62 AspleuAsplyslyslysThrValTrpArgleuProGluPheGlyGlnLeuIleLeuPhe
   Sequence 1344 BP; 326 A; 356 C; 390 G; 272 T; 0 U; 0 Other;
   1344
203
0
0
0
  US-10-048-116B-2_COPY_1_278 (1-278) x AAT60705 (1-1344)
   Length:
Matches:
Conservative:
Mismatches:
  Indels:
  Example 3; Page 129-132; 142pp; English.
   1.07e-100
1105.00
100.0%
100.0%
73.9%
  WPI; 1997-052337/05.
P-PSDB; AAW10513.
   Percent Similarity:
Best Local Similarity:
Query Match:
DB:
  Alignment Scores:
   102
  82
   142
   748
   122
   42
  ..
92
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Length:
Matches:
Conservative:
Mismatches:
Indels:

2.34e-98 1083.00 68.9% 65.0%

> Percent Similarity: Best Local Similarity:

Query Match: DB:

Alignment Scores: Pred. No.:

Score:

-----

```
867
  201
  927
   987
  ACCAGCTTCCTCGTCAACCGTGACCATTCCTTCCACAAGCTGTCTTATCTCACCTTCATC
  ProSerAspAspIleTyrAspCysLysValGluHisTrpGlyLeuGluGluBroVal
ThrSerPheLeuValAsnArgAspHisSerPheHisLysLeuSerTyrLeuThrPheIle
  ccrrcrargargacarrrangacrecaaggregagcacregegeceregaggaggecegerr
   Mouse; MHC; major histocompatibility complex; MHC class II; multimer; single chain; immunosuppressive; antidiabetic; antidiflammatory; antianaemic; antiarheritic; neuroproecetive; vaccine; autoimmune disease; insulin dependent diabetes; multiple sclerosis; myasthenia gravis; pernicious anaemia; autoimmune encephalomyelitis; rheumatoid arthritis; systemic lupus erythematosus; ss.
   Ä
   ŝ
  ABI99041 standard; cDNA; 1676
  Murine pCB223 coding sequence.
  22-MAR-2000; 2000US-0191274P.
15-MAY-2000; 2000US-0204249P.
23-JAN-2001; 2001US-0264003P.
  22-MAR-2001; 2001WO-US009616
   Carter D, Zhu S, Arimilli
   (first entry)
   GlyGlySer 224
   988 GGTGGCTCA 996
  WPI; 2001-616371/71.
P-PSDB; ABB56471.
   (CORI-) CORIXA CORP
   WO200170245-A1
   25-FBB-2002
  27-SEP-2001
   Mus sp.
Synthetic.
  182
  868
  808
   222
   ABI99041;
  3ULT 5
```

Multimeric complex for treating autoimmune diseases, comprises first and second single chain MrC class II molecules, each comprising alphai and betal domain linked through amino acid linker and multimerization domain.

Disclosure; Page 115; 147pp; English.

The invention relates to a multimeric complex comprising a first recombinant single chain major histocompatibility complex (MHC) class II molecule and a second recombinant single chain MHC class II molecule, and a second and a petal domain. The first and the second molecule are linked through the multimerisation domain. The first and the second molecule are linked through the multimerisation domain to form a multimeric complex, The complex is useful for treating autoimmune diseases. It is useful for treating insulin dependent diabetes, multiple sclerosis, myasthenia gravis, pernicious anaemia, autoimmune encephalomyelitis (EAE), rheumatoid arthritis and systemic lupus erythematosus. The present sequence encodes a single chain MHC class II molecule of the invention

Sequence 1676 BP; 438 A; 470 C; 407 G; 361 T; 0 U; 0 Other;

```
144
   456
  AsnileThrTrpLeuArgAsnSerLysSerValThrAspGlyValTyrGluThrSerPhe 164
  184
   204
   97 gacgacarrigaggccgaccacgraggcgrcrarggracaacrgrararcagrcrccrgga 156
  AsnPheThrProAlaThrAsnGluAlaProGlnAlaThrValPheProLysSerProVal 124
  517 crigicaaccereaccaricericeacaacciereriarereaccirearecerereac 576
   577 GATGATATTTATGACTGCAAGGTGGAGCACTGGGGCCTGGAGGAGCAGCCGGTTCTGAAACAC 636
  -----AlaGlnLysGluLeuGluLysGluLeuGln 237
  802 TCTGGATCCCTGTCCAGTGTGTGCACACCTTCCCAGCTGTCCTGCAGTCTGACCTCTAC 861
   238 AlaLeuGluLysGlu-----AsnAlaGlnLeuGluTrpGluLeuGlnAlaLeuGluLys 255
   TrpGluProGluIleProAlaProMetSerGluLeuThrGluThrGlyGlyGlyGlySer 224
  44
  64
  84
  277 GGGGACTGCAAAACATAGCTACAGGAAAATACACCTTGGGAATCTTGACTAAGGTGCA
   337 AATTCCACCCCAGCTACCAATGAGGCTCCTCAAGCGACTGTGTTCCCCCAAGTCCCCTGTG
   LeuLeuGlyGlnProAsnThrLeuIleCysPheValAspAsnIlePheProProValIle
   LeuValasnargaspHisSerPheHisLysLeuSerTyrLeuThrPheIleProSerAsp
  AspaspileTyrAspCysLysValGluHisTrpGlyLeuGluGluProValLeuLysHis
   256 GluLeuAlaGlnAlaAlaSer-----------GluProArgGlyPro
   157 gacarrescederacacacarearrerresresresresresresrrerrereser
  LysLysLysThrValTrpArgLeuProGluPheGlyGlnLeuIleLeuPheGluProGln
  217 AAGAAGGAGACTATCTGGATGCTTCCTGAGTTTGGCCAATTGACAAGCTTTGACCCCAA
   85 GlyGlyLeuGlnAsnIleAlaAlaGluLysHisAsnLeuGlyIleLeuThrLysArgSer
  457 AACATCACATGGCTCAGAAATAGTAAGTCAGTCACAGACGGCGTTTATGAGACCAGCTTC
   Achacaccccarrecercresecenteresecentereses and action of the contraction   742 GTGACTCTAGGATGCCTGGTCAAGGGTTATTTCCCTGAGCCAGTGACCTTGACCTGGAAC
   LeuIleLeuGlyValLeuAlaLeuAsnThrMetLeuSerLeuCysGly-----GlyGlu
   AspAspIleGluAlaAspHisValGlyPheTyrGlyThrThrValTyrGlnSerProGly
  AspileGlyGlnTyrThrHisGluPheAspGlyAspGluLeuPheTyrValAspLeuAsp
US-10-048-116B-2_COPY_1_278 (1-278) x ABI99041 (1-1676)
  ThrThrAlaProSer------
   105
  125
  145
  225
   25
  45
  65
   165
   185
   205
  229
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80

9

365

485

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66 TGTGGAGGTGAAGACGACATTGAGGCCGACCACGTAGGCACCTATGGTATAAGTGTATAT 125
   ThrLysArgSerAsnPheThrProAlaThrAsnGluAlaProGlnAlaThrValPhePro 120
  ProProValileAsnileThrTrpLeuArgAsnSerLysSerValThrAspGlyValTyr 160
   GluThrSerPheLeuValAsnArgAspHisSerPheHisLysLeuSerTyrLeuThrPhe 180
  181 ileProSerAspAsplleTyrAspCysLysValGluHisTrpGlyLeuGluGluPro 200
   ValleulyshisTrpGluEroGluIleProAlaProMetSerGluLeuThrGluThr 219
6 ATGCCGCGCAGCAGAGCTCTGATTCTGGGGGTCCTCGCCCTGACCACCATGCTCAGCCTC
  41 GlnSerProGlyAsp11eGlyGlnTyrThrHisGluPheAspGlyAspGluLeuPheTyr
  61 ValAspLeuAspLysLysLysThrValTrpArgLeuProGluPheGlyGlnLeuIleLeu
  CysGlyGlyGlyGluAspAspIleGluAlaAspHisValGlyPheTyrGlyThrThrValTyr
  PheGluProGlnGlyGlyLeuGlnAenIleAlaAlaGluLy8H$A8snLeuGlyIleLeu
   WO2005014849-A2.
  05-MAY-2005
  486
   81
  101
  161
   141
  201
   186
  909
  ADX26089
   Mus
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  g
  유
   쉽
  셤
   셤
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   요 요
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                                    ò
  The patent claims complexes of formulae (1), (11) and (111) which are as follows: (1) x - MCH - peptide; (11) MHC - peptide - x; (111) MHC - peptide. Wit (111) MHC - peptide. Wit (111) MHC - major histocompatibility glycoprotein; and the peptide includes an epitope associated with one of the major autoimmune diseases. The MHC is complexes can be used to treat and monitor the autoimmune diseases. The MHC is protocol, an oligonucleotide encoding the ACHR peptide 195-212 - an epitope in myasthenia gravis patients - was attached to the DNA encoding the N-terminal of the I-Ab-alpha chain. (Updated on 31-OCT-2002 to add missing OS field.) (Updated on 25-MAR-2003 to correct PR field.) (Updated
   I-A (Class II) histocompatibility protein;
major histocompatibility complex antigen; MHC-II; acetylcholine receptor;
myasthenia gravis; autoantigen; autommune disease; epitope.
   New complexes of histo-compatible glyco.protein - with antigenic peptide(\theta) and label or toxin, used to target antigen specific T helper cells.
       922 AATGTGGCCCACCCGGCAAGCAGCACCAAGGTGGACAAGAAATTGAGCCCAGAGGGCCC
  Class II major
  U; 0 Other;
   776
201
5
13
0
   Length:
Matches:
Conservative:
Mismatches:
Indels:
  Sequence 776 BP; 177 A; 205 C; 199 G; 195 T; 0
   982 acaarcaacccrercrercearceaaarceea 1014
  the
  Sequence encoding the I-Ab-alpha chain of histocompatibility complex (MHC) antigens.
  to correct PI field.)
  BR;
  Figure 8; Page ?; 74pp; English.
  BP
  Clark
  89WO-US002784
  88US-00210594
89US-00367751
   2.56e-98
1078.00
94.1%
91.8%
  AAQ03169 standard; DNA; 776
  (first entry)
   (revised)
  Lerch BL,
   (BIOS-) BIOSPAN CORP.
  WPI; 1990-022384/03
  on 25-MAR-2003
  cent Similarity:
  Unidentified
   23-JUN-1988;
21-JUN-1989;
  23-JUN-1989;
   unment Scores:
   WO8912459-A.
   25-MAR-2003
31-OCT-2002
  23-AUG-1990
   28-DEC-1989
  Sharma SD,
   268
   AAQ03169;
```

```
Novel cell pain response detection method-related mouse gene SeqID435
  pain; animal disease model; expression; analgesic; antiaddictive; nootropic; anticonvulsant; vasotropic; neuroprotective; tranquilizer; antisfematic; antiarthritic; osteopathic; ophthalmological; antiinflammatory; antipruritic; dermatological; antiulcer; gastrointestinal-Gen.; nephrotropic; gynecological; hepatocropic; antiparkinsonian; neuropetic; laxative; gene therapy; neuropathic pain; Alzheimers disease; Parkinsons disease; motor neurone disease; Huntingtons disease; schizophrenia; gene; ds.
                             ВЪ
   06-JUL-2004; 2004WO-US023166.
   03-JUL-2003; 2003US-0485101P.
                             ADX26089 standard; DNA; 978
   (first entry)
  SA.
  (EURO-) EUROCELTIQUE
  17-FEB-2005.
```

20

MetProCysSerArgAlaLeuIleLeuGlyValLeuAlaLeuAsnThrMetLeuSerLeu |||||||

0-048-116B-2\_COPY\_1\_278 (1-278) x AAQ03169 (1-776)

Similarity:

: y Match:

Local

: No ::

Lavery DJ; Chiang LW, Χu Υ, J1 R, WPI; 2005-163258/17 Tong J, Jin G,

Detecting pain responses in a cell, useful in identifying potential therapeutic and diagnostic candidates for treating pain, by identifying genes that are differentially expressed in a model of neuropathic pain.

Example 1; SEQ ID NO 435; 173pp; English

This invention relates to a novel method of detecting a pain response in a cell which comparises determining the expression level in a test cell of at levels in an animal model of pain, where similar or identical expression level to a level in an animal model of pain, where similar or identical expression levels indicate a pain response in the test cell. The invention may be useful for the development of compounds with an analgeaic, antiaddictive, notiopathic, antiaddictive, notiopathic, antiaddictive, antiaddictive, notiopathic, ostopathic, antimadictive, antiaddictive, notiopathic, ostopathic, antimadictive, antimactory, antiprivitic, dermatclogical, antiinflammaticy, antiprivitic, ostopathic, antimalial—den. nephrotropic, gynecological, hepatotropic, antiparkinsonian, neuroleptic or laxative activity whilst the disclosed sequences may prove useful for gene therapy. The methods and compositions of the present invention are useful for identifying agonists and antagonists for the gene or gene products as potential neuropathic pain, nociceptive pain, chonic pain, inflammatory pain, inflammatory discase, osteooriated with cancer, and pain associated with rheumatic disease, and also for addiction, seizure, stroke, ischemia, a neurodegenerative, osteooriated with cancer, and pain associated with rheumatic disease, osteooriated with cancer, and pain associated with rheumatic disease, osteooriated with cancer, and pain associated with rheumatic disease, and disorder, inflammatory bowel disease, irritable bowel disease (IBB), irritable bowel disease (IBB), gastrosophageal reflux disease (IBB), irritable bowel disease, inflammatory bowel disease, inflammatory explain, functional disarrhea, burbulence, chonical dispandant disarrhea, burbulence, chonical disease, desphageal chonical disease, desphageal cridin, functional disarrhea, burbulence, chonical disease, and charting and disease, maniforminal pain, recurrent abdominal pain, functional disarrhea, burbulence, chonical disease, maniformatical sequence and schizopheni the invention analysed in the method of

Sequence 978 BP; 218 A; 263 C; 247 G; 250 T; 0 U; 0 Other;

| 978<br>203               |                                                  |
|--------------------------|--------------------------------------------------|
| Length:<br>Matches:      | Conservative<br>Mismatches:<br>Indels:<br>Gaps:  |
| 7.05e-98<br>1075.00      | 93.6%<br>92.7%<br>71.9%                          |
| ignment Scores: ad. No.: | cont Similarity: s: Local Similarity: ary Match: |

-10-048-116B-2\_COPY\_1\_278 (1-278) x ADX26089 (1-978)

- 83 24 ATGCCGCCGCAGCAGAGCTCTGATTCTGGGGGTCCTCGCCCTGACCACCATGCTCAGCCTC 1 MetProCysSerArgAlaLeuIleLeuGlyValLeuAlaLeuAsnThrMetLeuSerLeu
- 40 21 CysGlyGlyGluAspAspIleGluAlaAspHisValGlyPheTyrGlyThrValTyr
- 9 41 GlnSerProGlyAspIleGlyGlnTyrThrHisGluPheAspGlyAspGluLeuPheTyr
- ValAspLeuAspLysLysLysLysThrValTrpArgLeuProGluPheGlyGlnLeuIleLeu 80 61

| 97 81 PheGluprocalcollogical controlled by PheGluprocalcollogy High High High High High High High High | arma SD, Lerch BL, Clark  1; 1993-036056/04.  re major MHC-peptide complesponse such as auto:immunit | seguence (AAT0628<br>nstruction of nov<br>rived e.g. from t |
|--------------------------------------------------------------------------------------------------------|------------------------------------------------------------------------------------------------------|-------------------------------------------------------------|
|--------------------------------------------------------------------------------------------------------|------------------------------------------------------------------------------------------------------|-------------------------------------------------------------|

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120
   140
   425
   | IleProSerhspAspAspIleTyrAspCysLysValGluHisTrpGlyLeuGluGluPro 200
  605
   66 TGTGGAGGTGAAGACGACATTGAGGCCGACCAGTAGGCACCTATGGTATAAGTGTATAT 125
   MHC; major histocompatibility complex; toxic conjugate; autoimmunity; autoimmune disease; helper T-cell; T-lymphocyte; acetylcholine receptor; myelin basic protein; I-Ab; ds.
   9
  ValAapLeuAapLyaLyaLyaThrValTrpArgLeuProGluPheGlyGlnLeuIleLeu 80
   20
   65
   40
appropriate MHC molecule via the I-Ab-alpha (or -beta) chain sequence. The resulting conjugate is used in the treatment of autoimmune diseases
   ValLeuLysHisTrpGluProGluIleProAlaProMetSerGluLeuThrGluThr 219
  ThrLygArgSerAsnPheThrProAlaThrAsnGluAlaProGluAlaThrValPhePro
  GlnSerProGlyAsp11eGlyGlnTyrThrHisGluPheAspGlyAspGluLeuPheTyr
   306 ACTAAGAGGTCAAATTCCACCCCAGCTACCAATGAGGCTCCTCAAGGGACTGTTCCCC
   141 ProprovallleAsnlleThrTrpLeuArgAsnSerLysSerValThrAspGlyValTyr
  426 CCTCCTGTGATCAACATCACATGGCTCAGAAARAGCAAGTCAGTCGCAGACGGTGTTTAT
  GluThrSerPheLeuValAsnArgAspHisSerPheHisLysLeuSerTyrLeuThrPhe
  CysGlyGlyGluAspAspIleGluAlaAspHisValGlyPheTyrGlyThrThrValTyr
  LysSerProValLeuLeuGlyGlnProAsnThrLeuIleCysPheValAspAsnIlePhe
  Sequence 776 BP; 177 A; 205 C; 199 G; 194 T; 0 U; 1 Other;
  776
200
5
14
0
  Length:
Matches:
Conservative:
Mismatches:
Indels:
Gaps:
   0-048-116B-2_COPY_1_278 (1-278) x AAT06285 (1-776)
   1.02e-97
1072.00
93.6%
91.3%
   AAQ56919 standard; DNA; 776
   (revised)
(first entry)
   Mouse I-Ab-alpha chain.
   ent Similarity:
   25-MAR-2003
08-JUL-1994
   unment Scores:
  101
  121
  201
   486
  AAQ56919;
  21
   41
   126
   61
   186
  81
   191
   181
  y Match:
   ٠٠ No.:
```

```
The acetylcholine receptor alpha subunit given in sequence AAR45946 (corresponding mRNA in AAQ56918) and the myelin basic protein given in AAA45947 are antigenic peptides associated with autoantigens. They have been conjugated with MHC class II components such as I-Ab- alpha chain (encoded by sequene AAQ56919) and I-Ab-beta chain, (encoded by AAQ56920) and a toxin or label to form conjugates used to target helper T-cells for the treatment of autoimmune diseases. (Updated on 25-MAR-2003 to correct PF field.)
  CAGTCTCCTGGAGACATTGGCCAGTACACATTTGAATTTGATGCTGATGATTGTTCTAT
  ThriysArgSerAsnPheThrProAlaThrAsnGluAlaFroGlnAlaThrValPhePro
  LysSerProValLeuleuGlyGlnProAsnThrLeulleCysPheValAspAsnIlePhe
   CysG1yG1yG1ydspAspI1eG1uAlaAspHisValG1yPheTyrG1yThrThrValTyr
  VallaspleulasplyslyslyslyslhrvalfrplargleuproglupheglyglnLeulleLeu
   PheGluProGlnGlyGlyLeuGlnAenIleAlaAlaGluLysHisAsnLeuGlyIleLeu
  GlaserProGlyAspIleGlyGlnTyrThrHisGluPheAspGlyAspGluLeuPheTyr
  Met ProCysSerArgAlaLeu1leLeuGlyValLeuAlaLeuAsnThrMetLeuSerLeu
  MHC-mediated toxic peptide conjugates - useful for ameliorating auto-
   BP; 177 A; 205 C; 199 G; 194 T; 0 U; 1 Other;
   Length:
Matches:
Conservative:
Mismatches:
Indels:
   US-10-048-116B-2_COPY_1_278 (1-278) x AAQ56919 (1-776)
   Disclosure, Fig 8.1-8.2; 42pp; English
   BR;
   88US-00210594.
89US-00367751.
90US-00576084.
   Clark
  90US-00635840
   1.02e-97
1072.00
93.6%
91.3%
   Lerch BL,
   WPI; 1994-056406/07.
   Score:
Percent Similarity:
Best Local Similarity:
Query Match:
DB:
   (ANER-) ANERGEN INC.
   P-PSDB; AAR45948
  Sequence 776
   Alignment Scores:
Pred. No.:
  28-DEC-1990;
                                  US5284935-A.
  23-JUN-1988;
   30-AUG-1990
  08-FEB-1994
   SD,
  immunity.
  9
  126
   186
   306
   21
  99
   41
   61
  81
   246
  101
  121
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80

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365

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WO9604314-A1

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  ð
                      426 CCTCCTGTGATCAACATCACATGGCTCAGAAARAGCAAGTCAGTCGCGCGGGGTGTTAT 485
  /label= I-Ad alpha-TM
/note= "murine MHC class II I-Ad gene alpha-transmembrane
domain"
ProProvallleAsnIleThrTrpLeuArgAsnSerLysSerValThrAspGlyValTyr 160
  161 GluThrSerPheLeuValAsnArgAspHisSerPheHisLysLeuSerTyrLeuThrPhe 180
   181 IleProSerAspAspAspIleTyrAspCysLysValGluHisTrpGlyLeuGluGluPro 200
  546 ATCCCTTCTGACGATGACATTTATGACTGCAAGGTGGAACACTGGGGCCTGGAGGAGCCG 605
  6. 1508
/*tag= a
6. 86
/*tag= b
/*tag= b
/*tag= b
/label= I-Ad_beta_chain_leader
/note= "murine MHC class II I-Ad gene beta chain leader
  GTTCTGAAACACTGGGAACCTGAGATTCCAGCCCCCATGTCAGAGACTGACAGAGACT 662
   201 ValleuLysHisTrpGluProGluIleProAlaProMetSerGluLeuThrGluThr 219
   MHC; major histocompatibility complex; PCR; polymerase chain reaction; T cell activity modulator; antagonist; immune disorder; allergy; multiple aclerosis; insulin-dependent diabetes mellitus; rheumatoid arthritis; myasthenia gravis; ds.
   /*tag= h
/label= I-Ad_alpha1
/note= "murine MHC class II I-Ad gene alpha-2 domain"
1068. .1352
   /*tag= 1
/label= I-Ad_alpha2
/note= "murine MHC class II I-Ad gene alpha-2 domain"
1353. .1505
   /*tag= e .
//label= I-Ad_betal
//label= I-Ad_betal
53. .734
//rag= f .
//label= I-Ad_beta2
//note= "murine MHC class II I-Ad gene beta-2 domain"
  Vector SCT1-derived single chain gene encoding MHC fusion complex.
  **tag= c
/label= OVA 323-339
//note= "chicken ovalbumin residues 323-339"
138. .167
   *tag= d
note= "10 residue linker peptide"
68. .452
   /*tag= g
/note= "24 residue peptide linker"
307. .1067
   Location/Qualifiers
  AAT17587 standard; DNA; 1508 BP
   26-SEP-1996 (first entry)
  "eodnence"
  35. .806
  misc feature
  misc_feature
  misc_feature
   misc_feature
   misc_feature
  misc_feature
   misc_feature
   misc_feature
   sig_peptide
   Synthetic.
   AAT17587;
   486
  909
   Key
CDS
```

```
AAT17587 encodes a murine MHC fusion complex capable of modulating T cell activity encoded by the vector SCT1. The MHC fusion complex comprises at least one MHC molecule containing a peptide-binding groove and a presenting peptide covalently linked to the MHC molecule and opt. a transmembrane domain. DNA encoding a MHC fusion complex may be cloned into a host cell to express the complex. The transformed cells may then complex to the used to identify peptides that modulate, pref. antegonise, T cell activity. DNA encoding a MHC fusion complex or a single chain fusion complexes may be used to suppress an immune response in an animal suffering from an immune disorder e.g. multiple sclerosis, insulin dependent diabetes mellitus, rheumatoid arthritis, myasthemia gravis or chronic allergies. The complexes may also be used in the treatment of chronic allergies. The complexes may also be used in the treatment of chronic allergies and dogs. The MHC fusion complexes can be produced such that they contain a single antigenic peptide including one of known structure, additionally a wide range of peptides can be presented for T cell interaction
  912 TTCTATGTGGACTTGGATAAGAAAACTGTCTGGAGGCTTCCTGAGTTTGGCCAATTG 971
   792 decegrirecrecagneaagaegaearingaeseceaecaeeraeserieraresraeaaer 851
  59 PheTyrValAspLeuAspLysLysLysThrValTrpArgLeuProGluPheGlyGlnLeu 78
   22 GlyGly------GluAspAspIleGluAlaAspHisValGlyPheTyrGlyThrThr
  Major histocompatability complex fusion complex for modulating T ce. activity - used in the treatment of immune disorders, e.g. multiple sclerosis, IDDM and rheumatoid arthritis.
  Sequence 1508 BP; 337 A; 414 C; 440 G; 317 T; 0 U; 0 Other;
   Edwards AC;
  0064
   US-10-048-116B-2_COPY_1_278 (1-278) x AAT17587 (1-1508)
   Length:
Matches:
Conservative:
Mismatches:
Indels:
   Grammer S,
  Example 17; Fig 28; 210pp; English.
   Weidanz JA,
  95WO-US009816.
  94US-00283302
95US-00382454
   1.15e-96
1065.50
98.5%
98.5%
71.2%
   Chavaillaz P, Jiao J;
   (DADE-) DADE INT INC.
   Rhode PR,
   WPI; 1996-129343/13.
   Percent Similarity:
Best Local Similarity:
  P-PSDB; AAR98906
  31-JUL-1995;
  29-JUL-1994;
01-FEB-1995;
   Alignment Scores:
  15-FEB-1996.
   Wong HC,
  Query Match:
   Score:
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79 IleLeuPheGluProGlnGlyGlyLeuGlnAsnIleAlaAlaGluLysHisAsnLeuGly

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ATACTCTTTGAGCCCCAAGGTGGACTGCAAAACATAGCTGCAGAAAAACACAACTTGGGA 1031
  1032 ATCTTGACTAAGAGGTCAAATTTCACCCCAGCTACCAATGAGGCTCCTCAAGCGACTGTG 1091
   VallyrGluThrSerPheLeuValAsnArgAapHisSerPheHisLysLeuSerTyrLeu 178
  IlePheProProValileAsnileThrTrpLeuArgAsnSerLysSerValThrAspGly 158
   Major histocompatibility complex; MHC; single chain MHC; sc-MHC; Ig; peptide binding groove; immunoglobulin; T cell receptor; immune response; immune-related disorder; antigenic peptide; fusion protein; ss.
  PheTyrValAspLeuAspLysLysLysLysThrValTrpArgLeuProGluPheGlyGlnLeu
  TTCTATGTGGACTTGGATAAGAAGAAAACTGTCTGGAGGCTTCCTGAGTTTGGCCAATTG
   ThrPheIleProSerAspAspAspIleTyrAspCysLysValGluHisTrpGlyLeuGlu
  ACCTTCATCCCTTCTGATGATGACATTATGACTGCAAGGTGGAGCACTGGGGCCTGGAG
  GlyGly------GluAspAspIleGluAlaAspHisValGlyPheTyzGlyThzThz
|||||||
|GCGGTTCCTCGAGTGAAGACGACTATGAGGCCGACCACGTAGGCTTCTATGGTACAACT
  ValTyrGinSerProGlYAspileGlYGlnTyrThrHisGluPheAspGlyAspGluLeu
  | IleLeuPheGluProGlnGlyGlyLeuGlnAsnIleAlaAlaGluLysHisAsnLeuGly
   Phe ProlysSer ProValLeuLeuGlyGlnProAsnThrLeuIleCysPheValAspAsn
  IleLeuThrLysArgSerAsnPheThrProAlaThrAsnGluAlaProGlnAlaThrVal
   Single chain IAd/OVA 323-229 MHC fusion protein encoding DNA
  US-10-048-116B-2_COPY_1_278 (1-278) x AAT86988 (1-1508)
Matches:
Conservative:
Mismatches:
  Indels:
   BP
   AAX89069 standard; DNA; 1508
   98WO-US021520
1065.50
98.5%
98.5%
71.2%
  (first entry)
            Percent Similarity:
Best Local Similarity:
Query Match:
DB:
   ACT 1394
  Thr 219
  14-SEP-1999
   13-OCT-1998;
  WO9921572-A1
  06-MAY-1999,
  Synthetic
  1092
   139
   159
   1212
   179
   1272
  219
  792
  39
   852
  912
   79
   972
  66
   1392
  22
   59
  119
   AAX89069;
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  셤
   ઠે
  영 장
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   1331
  1151
   Single chain major histocompatibility complex comprising linked alpha and beta chains - useful for suppressing an immune response to an auto:immune disease, e.g. multiple sclerosis, rheumatoid arthritis, diabetes mellitus, etc.
                138
  178
  158
  179 ThrPhelleProSerAspAspAspIleTyrAspCysLysValGluHisTrpGlyLeuGlu 198
                159 ValTyrGluThrSerPheLeuValAsnArgAspHisSerPheHisLysLeuSerTyrLeu
   1272 ACCTTCATCCCTTCTGATGATGACATTTATGACTGCAAGGTGGAGCACTGGGGCCTGGAG
  139 ilePheProProValIleAsnIleThrTrpLeuArgAsnSerLysSerValThrAspGly
  Construction; major histocompatibility complex; MHC; fusion complex; SCT1 single chain gene; 88.
  Sequence 1508 BP; 337 A; 413 C; 441 G; 317 T; 0 U; 0 Other;
  the construction of major
  1508
  fusion complexes
   Wong HC
  Length:
   Example 17; Page 137-139; 217pp; English.
  Location/Qualifiers
6. .1508
/*tag= a
  The present sequence was used in histocompatibility complex (MHC)
   Σ
   AAT86988 standard; DNA; 1508 BP
   Burkhardt
  97WO-US001617
   96US-00596387
  1.15e-96
  (first entry)
  SCT1 single chain gene
  (DADE-) DADE INT INC
   WPI; 1997-402555/37.
P-PSDB; AAW29213.
   Rhode PR, Jiao J,
   1392 ACT 1394
  219 Thr 219
   unment Scores:
  30-JAN-1997;
   31-JAN-1996;
   WO9728191-A1
  27-MAR-1998
   07-AUG-1997
  Synthetic
   AAT86988;
   Key
```

1211

118

851

911

28

971

98

78

138

1271

1331

```
Single chain major histocompatibility complex class I complexes
  Wong HC;
  Jiao J,
  Rhode PR, Acevedo J, Burkhardt M,
97US-00960190.
  (SUNO-) SUNOL MOLECULAR CORP.
  WPI; 1999-418411/35.
P-PSDB; AAY27111.
29-OCT-1997;
```

Example 1; Fig 1; 148pp; English.

The invention relates to new single chain major histocompatibility complex (sc-MHC) class II complexes that comprise a peptide binding groove, and a modified class II beta 2 chain or covalently linked immunoglobulin (Ig) light chain contant (II) region. The MHC complexes are useful for detection and analysis of peptide ligands, pathogenic Teals, for functional, cellular and molecular assays. They can be used to identify and isolate T cell receptor and/or MHC agonists and antagonists. They can also be used to calls involved in immune-related disorders. They can also be used to raise antibodies and to screen immune cells. It is also use in a method of suppressing an immune response in mammals. The sc-MHC complexes comprising modified class II beta 2 chains and/or Ig-C1 regions are soluble and provide enhanced yield. These MHC complexes also contain single antigent cpetides readily isolated from expressing cells in significant quantities. The polyspecific MHC complexes also provide a means to detect cells expressing multiple target structures with a single complex. The present sequence represents a DNA encoding a single chain IAd/OVA 323-229 MHC fusion protein

Sequence 1508 BP; 337 A; 413 C; 441 G; 317 T; 0 U; 0 Other;

| 1508<br>198<br>0 0<br>3                                          | 508)                  | GlyGlyGluAspAsp1leGluAlaAspHisValGlyPheTyrGlyThrThr 38<br>      <br>GGCGGTTCCTCGAGTGAAGACGACTATGAGGCCGACCACGTAGGCTTCTATGGTACAACT 851 | ValTyrginserproglyaspilegiyginTyrThrHisGluPheAspGlyAspGluLeu 58<br> | Phetyrvalaspleuasplyslyslysthrvaltrpargleuproglupheglyglnleu 78<br> | IleleuPheGluProGlnGlyGlyLeuGlnAsnIleAlaAlaGluLysHisAsnLeuGly 98<br> |                                                            | PheProLysSerProValLeuLeuGlyGlnBroAsnThrLeuIleCysPheValAspAsn 138<br> | IlePheProProValileAsnileThrTrpLewargAsnSerLysSerValThrAspGly 158<br> |
|------------------------------------------------------------------|-----------------------|--------------------------------------------------------------------------------------------------------------------------------------|---------------------------------------------------------------------|---------------------------------------------------------------------|---------------------------------------------------------------------|------------------------------------------------------------|----------------------------------------------------------------------|----------------------------------------------------------------------|
| Length: Matches: Conservative: Mismatches: Indels: Gaps:         | x AAX89069 (1-1508    | AspileGlualaAspH<br>            <br>:GACATTGAGGCCGACC                                                                                | ileglyglnTyrThrH<br>                 <br> arrggccagracacac          | LysLysThrvalTrpA:<br>                                               | GlyLeuGlnAsnIleA<br>                                                | PhethrProAlathra<br>                                       | iLeuglyglnProAsnT<br>                                                |                                                                      |
| 1.15e-96<br>1065.50<br>1065.5<br>98.5<br>1ty: 98.5<br>2          | OPY_1_278 (1-278)     | 1yGluasp<br>  <br>                                                                                                                   | yrGlnSerProGlyAsp<br>                                               | yrvalaspleuasplys<br>                                               | euPheGluProGlnGly<br>                                               | euthrLysArgSerAsr<br>                <br>TGACTAAGAGGTCAAAT | roLysSerProValleu<br>                                                | heProProVallleAsn<br>                <br>rcccaccrGrGarcaac           |
| ignment Scores: ed. No.: Dre: cent Similarity: st Local Similari | -10-048-116B-2_COPY_1 | 22 GlyGly<br>       <br>792 GGCGGT                                                                                                   | 39 Vall<br>    <br>1852 GTTT                                        | 59 PheT)<br>     <br>912 TTCT                                       | 79 IleL6<br>     <br> 972 ATAC                                      | 99 IleL<br>    <br>1032 ATCT                               | 119 PheP:<br>    <br> 1                                              | 139 IleP<br>    <br> 1152 ATCT                                       |

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1331
  The invention relates to a major histocompatibility complex (WHC) fusion complex (I) comprising an MHC molecule that contains a peptide-binding groove, and a presenting peptide covalently (e.g. an antiganic peptide) linked to the MHC molecule, where (I) is capable of modulating the activity of a T cell. Also included are a DNA construct coding for the complex, where the MHC molecule is a class II MHC (e.g. mouse I.Ad or I-As or human HLA-DR1 (human leukocyte antigen-DR1), a multivalent MHC fusion complex compraints to or more linked complexes, identifying a peptide that can modulate the activity of T cells (involving introducing into host cells cloning vectors that each contain the fusion complex DNA, culturing the host cells under conditions suitable for expression of the MHC fusion complex, and selecting host cells that express MHC fusion complex that modulate the activity of T cells), a single recombinant
  Novel major histocompatibility complex fusion complex having presenting peptide covalently linked to MHC molecule containing peptide-binding groove, used for suppressing immune response in multiple sclerosis,
   179 ThrPheIleProSerAspAspAspIleTyrAspCysLysValGluHisTrpGlyLeuGlu
  1272 ACCTTCATCCCTTCTGATGATGACATTTATGACTGCAAGGTGGAGCACTGGGGCCTGGAG
ValTyrGluThrSerPheLeuValAsnArgAspHisSerPheHisLysLeuSerTyrLeu
  MHC; major histocompatibility complex; gene therapy; fusion complex peptide-binding groove; T cell modulation; class II MHC; vaccine; autofimmune disorder; multiple sclerosis; rheumatoid arthritis; insulin-dependent diabetes mellitus; myasthenia gravis; immunogen; chronic allergy; mouse; ds; I-Ad; gene.
  Edwards AC;
  Mouse MHC I-Ad/Ova 323-339 synthetic gene SCT1.
  o,
  Grammer
  Example 17; Fig 28; 126pp; English.
  Rhode PR, Weidanz JA, z P, Jiao JJJ;
  ACA60743 standard; DNA; 1508 BP.
  94US-00283302.
95US-00382454.
97US-00776084.
   06-JUL-2001; 2001US-00900379.
  (first entry)
   (DADE-) DADE INT INC.
   WPI; 2003-341126/32.
P-PSDB; ABU72107.
   1392 ACT 1394
  219 Thr 219
  US2002198144-A1.
  Chavaillaz P,
  29-JUL-1994;
01-FEB-1995;
  16-JUN-2003
  17-JAN-1997;
  26-DEC-2002.
   Mus sp.
Synthetic.
   allergies.
   ACA60743;
  Wong HC,
  RESULT 13
  ACA60743
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expression vector comprising DNA that codes for the alpha and beta chains of the fusion complex MHC protein, a single recombinant expression vector comprising DNA that codes for a T cell costimulatory factor and the alpha and beta chains of the MHC fusion complex. The DNA constructs can contain heterologous leader peptide sequences and Kozak sequence for efficient expression of the fusion complex. Also included are inducing an immune response in a mammal (including vaccinating a mammal against a targeted disorder, by administering DNA sequence comprising a fusion complex, or DNA sequence coding for a fusion complex which is a single chain fusion molecule) and suppressing an immune response in a mammal by administering to the mammal a DNA sequence comprising an expression vector, encoding a full length MHC molecule that contains a transmembrane domain, and a presenting peptide that is a T cell receptor (TCP) antagonist or partial agonist and is covalently linked to the MHC protein, or DNA sequence coding for the fusion complex which is a single chain fusion molecule. The methods are useful for identifying a peptide that can modulate the activity of T cells, inducing an immune response in a mammal against a targeted disorder) and for suppressing an immune response in a mammal. The disorders include an autoimmune disorder such as multiple sclerosis, insulin-dependent diabetes mallitus, rheumatoid arthritis, myasthenia gravis or chronic allergies. The present sequence encodes a mouse MHC class II I-Ad fusion complex of the

Sequence 1508 BP; 337 A; 413 C; 441 G; 317 T; 0 U; 0 Other;

```
1152 Arctroccacororidaticaacarcacargecroagaaaragcaagroagroacagacgec 1211
   1272 ACCTTCATCCCTTCTGATGATGACATTTATGACTGCAAGGTGGAGCACTGGGGCCTGGAG 1331
   1092 frecedaherecerrerecresereserekeereka 1151
   118
  PheProLysSerProValLeuLeuGlyGlnProAsnThrLeuIleCysPheValAspAsn 138
   139 IlePheProProValileAsnileThrTrpLeuArgAsnSerLysSerValThrAspGly 158
  159 VallyrdluThrSerPheLeuValAsnArgAspHisSerPheHisLysLeuSerTyrLeu 178
   179 ThrPhelleProSerAspAspAspIleTyrAspCysLysValGluHisTrpGlyLeuGlu 198
  851
   852 GTTTATCAGTCTCCTGGAGACATTGGCCAGTACACACATGAATTTGATGGTGATGAGTTG 911
  98
  28
   PheTyrValAspLeuAspLysLysLysLysLysLysThrValTrpArgLeuProGluPheGlyGlnLeu 78
  39 ValtyrGlnSerProGlyAspileGlyGlnTyrThrHisGluPheAspGlyAspGluLeu
  79 IleLeuPheGluProGlnGlyGlyLeuGlnAsnIleAlaAlaGluLysHisAsnLeuGly
   99 IleLeuThrLysArgSerAsnPheThrProAlaThrAsnGluAlaProGlnAlaThrVal
   GlyGly------GluAspAspIleGluAlaAspHisValGlyPheTyrGlyThrThr
   0-048-116B-2_COPY_1_278 (1-278) x ACA60743 (1-1508)
  Length:
Matches:
Conservative:
Mismatches:
Indels:
  Gaps:
1.15e-96
1065.50
98.5%
98.5%
71.2%
   ent Similarity:
Local Similarity:
  119
   29
```

199 GluProValLeuLysHisTrpGluProGluIleProAlaProMetSerGluLeuThrGlu 218

```
A novel soluble fused major histocompatibility complex (MHC)

heterodimer:peptide complex, comprises DNA encoding 1st and 2nd MHC

comains, e.g. the present sequence, linked by DNA encoding a 5-25 residue

linker, and a DNA encoding an antiquenc peptide able to associate with a

copplide binding groove of the MHC molecule, linked in frame to the DNA

complex can be used to induce immunological tolerance in adults

complex can be used to induce immunological tolerance in adults

cusceptible to, or suffering from an autoantigen related disease, e.g.

susceptible to, or suffering from an autoantigen related disease, e.g.

cusceptible to, or suffering from an autoantigen related disease, e.g.

insulin dependent diabetes mellitus (IDDM), by antagonising the binding

of particular T cells and antigen presenting cells, to induce anergy

(immunological non-responding antigen are permanently linked into a

single chain, obviating the requirement for complex heterodimer

truncation or formation, the complex eliminates inefficient and non-

specific peptide loading. (Updated on 27-AUG-2003 to correct OS field.)
1332 GAGCCGGTTCTGAAACACTGGGAACCTGAGATTCCAGCCCCCATGTCAGAGCTGACGAA 1391
  Novel fused major histocompatibility complex:antigenic peptide complex - useful to induce tolerance to an autoantigen-related disease e.g. insulin-dependent diabetes mellitus.
   Soluble; fusion; major histocompatibility complex; MHC; region; heterodimer; complex; alpha2; antigen, binding groove; tolerance; autoantigen; disease; insulin dependent; diabetes mellitus; IDDM; antagonist; T cell; anergy; presenting cell; NOD mouse; Class II; alpha1;
  Sheppard PO;
  Seguence 588 BP; 153 A; 153 C; 136 G; 146 T; 0 U; 0 Other;
  Alphalalpha2 region of Class II NOD mouse MHC (IAg7) cDNA.
  Gross JA, Deshpande S,
  Example 3; Page 132-133; 142pp; English.
   Location/Qualifiers
1. .588
/*tag= a
   AAT60698 standard; cDNA; 588 BP.
   95US-00480002.
95US-00482133.
95US-00483241.
95US-0005964P.
   96WO-US010102
  (revised)
(first entry)
  EP,
  INC.
  (ZYMO ) ZYMOGENETICS (ANER-) ANERGEN INC.
  Kindsvogel W, Reich
   WPI; 1997-052337/05.
P-PSDB; AAW10505.
   1392 ACT 1394
  219 Thr 219
   07-JUN-1996;
  WO9640944-A2
  27-AUG-2003
12-SEP-1997
  07-JUN-1995
   07-JUN-1995
27-OCT-1995
  mat_peptide
  19-DEC-1996
   07-JUN-1995
   AAT60698;
   вр.
   Mus
  AAT60698
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92US-00869293

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  163
  420
   480
   120
  180
   103
  123
   143
   183
  203
   9
   63
  83
   61 GGAGACATTGGCCAGTACACACATGAATTTGATGGTGATGAGTTGTTGTTGTGGACTTG
   GlnGlyGlyLeuGlnAsnIleAlaAlaGluLysHisAsnLeuGlyIleLeuThrLysArg
   SerAsnPheThrProAlaThrAsnGluAlaProGlnAlaThrValPheProLysSerPro
  301 GIGCIGCIGGGICAGCCCAACACCCCTAICTGCTTIGIGGACAACAICTICCCACCTGIG
  144 IleAsnIleThrTrpLeuArgAsnSerLysSerValThrAspGlyValTyrGluThrSer
   361 ATCAACATCACATGGCTCAGAAATAGCAAGTCAGTCACACAGACGGCGTTTATGAGACCAGC
  AspAspAspIleTyrAspCysLysValGluHisTrpGlyLeuGluGluProValLeuLys
   GATGATGACATTTATGACTGCAAGGTGGACACTGGGGGCCTGGAGGAGCCGGTTCTGAAA
   GlyapileGlyGlnTyrThrHisGluPheAspGlyAspGluLeuPheTyrValAspLeu
   64 AsplyslyslyslystyaltrpArgleuProGluPheGlyGlnLeuIleLeuPheGluPro
  241 TCAAATTTCACCCCAGCTACCAATGAGGCTCCTCAAGCGACTGTGTTCCCCCAAGTCCCCT
   PheLeuValAsnArgAspHisSerPheHisLysLeuSerTyrLeuThrPhelleProSer
   GluAspAspIleGluAlaAspHisValGlyPheTyrGlyThrThrValTyrGlnSerPro
  1 GAAGACGACATTGAGGCCGACCACGTAGGCTTCTATGGTACAACTGTTTATCAGTCTCCT
  421 TTCCTCGTCAACCGTGACCATTCCTTCCACAAGCTGTCTTATCTCACCTTCATCCTTCT
   124 ValLeuLeuGlyGlnProAsnThrLeuIleCysPheValAspAsnIlePheProProVal
  receptor; epitope; myasthenia gravis; MG; antigen;
MHC; IAB; alpha; beta; ds.
   588
   HisTrpGluProGluIleProAlaProMetSerGluLeuThrGluThr 219
  588
1196
0
0
0
         Length:
Matches:
Conservative:
Mismatches:
Indels:
   AAT60698 (1-588)
   Gaps:
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The AChR peptide 195-215, which has been characterised as an epitope in myasthenia gravis (MG) in humans and in mice, may be connected to the Neterminal antigen binding site of a polypeptide derived from an MHC antigen associated with MG. For example, if the recombinant complex is to be used in mice, the AChR peptide may be incorporated into a sequence encoding either the I-Ab-alpha or I-Ab-beta chain (see AAQ35054-55 respectively). If the AChR peptide is to be incorporated into the beta chain, for example, the oligonucleotide may be inserted as a replacement for the leader sequence. (Updated on 25-MAR-2003 to correct PN field.)
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   9
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Mismatches:
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Recombinant proteins and molecular complexes derived therefrom, analogous to molecules involved in immune responses
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Location/Qualifiers

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Synthetic construct

synthetic construct

construct

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cher sequences; artificial sequences.

cher sequences; artificial sequences.

cher sequences; artificial sequences.

cher sequences; artificial sequence; bolyvalent WHC-binding domain fused proteins and conjugates, and utilization

polyvalent WHC-binding domain fused proteins and conjugates, and utilization

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polyvalent WHC-binding domain fused proteins and conjugates, and utilization

AL Patent: JP 2002504342-A/7

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PR 19-FEB-1999 UP 2000532537

PR 19-FEB-1999 US 60/075351

PR AJ W WUCHERPFENNIG 4/48, CO7K14/705, CO7K19/00, CC 10201/02,

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|  | 880<br>499<br>940                                                                                                                                                                                                                                                                                                                                                                                                                                                   |                                                                                                                                                                                                                                                                                    | 1120<br>739<br>1180<br>799 | QY         1240 ACCTGCATGGTCACAGACTTCATGCCTGAAGACATTACGTGGAGTGGACCAACAACGGG 1299                                                                                                   | 919 ADAACAGAGCTAAACTACAAGAACACTGAACCAGTCCTGGAACTCTGATGGTTCTTACTTC  1360 ATGTACAGCAAGCTGAGAAAAGAAAGAACTGGGTGGAAAGAAA | Qy         1420 TCAGTGGTCCACGAGGTCTGCACAATCACAACGAGTTTTCCCGGACTCGG 1479                                                                                                                                                                         | RESULT 5 CQ806532 LOCUS LOCUS CQ806532 LOCUS SEQUENCE 27 from Patent WO2004035622. ACCESSION CQ806532 LOI:47111926 VERSION CQ806532.1 GI:47111926 KEYWORDS SOURCE ORGANISM SYNTHetic construct ORGANISM REFERENCE AUTHORS Dreher I. and Moll.T. TITLE AUTHORS ALTEGRANISM 11-15 JOURNAL PATENT. WO 2004035622-A 27 29-APR-2004; F. HOFFMANN-LA ROCHE AG (CH)                                                                                     |
|--|---------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|----------------------------|------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|---------------------------------------------------------------------------------------------------------------------|-------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|--------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
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Sciurognathi, Muroidea, Muridae, Murinae, Mus.
  Direct Submission
Submitted (18-SEP-2000) McLean G.R., Cell Biology, Albert Eins
College of Medicine, 1300 Morris Park Avenue, Bronx, New York
  McLean, G.R., Nakouzi, A., Casadevall, A. and Green, N.S.
Human and murine immunoglobulin expression vector cassettes
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JOURNAL
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REFERENCE
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JOURNAL
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Best Local Similarity 100.0%; Pred. No. 1.8
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MANU294738 990 bp mRNA linear ROD 09-FEB-2001 Mus musculus partial mRNA for immunoglobulin heavy chain constant

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1 (bases 1 to 1341)
Beavers, L.S., Bumol, T.F., Gadski, R.A. and Novel recombinant and chimeric antibodies adenocarcinoma antigen
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  1 (bases 1 to 1095)
Sikorav,J.L., Auffray,C. and Rougeon,F.
Structure of the constant and 3' untranslated regions of the murine
Balb/c gamma 2a heavy chain messenger RNA
Nucleic Acids Res. 8 (14), 3143-3155 (1980)
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MOUSE mRNA for gamma-2a-immunoglobulin heavy-chain.
V00798
V00798.1 GI:51835
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Mus musculus
Musculus
Eukaryota, Metazoa, Chordata, Craniata, Vertebrata, Euteleostomi,
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Sciurognathi, Muroidea, Muridae, Murinae, Mus.
1 (bases 1 to 1407)
Lai, Y.S., John, J.A.C., Guo, I.C., Chen, S.C., Fang, K. and Chang, C.Y.
In vitro efficiency of intra- and extracellular immunization with
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M-csf-specific monoclonal antibody and uses thereof
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Lai,Y.-S., John,C., Guo,I.-C., Chen,S.-C., Fang,K. and Chang,C.-Y. Direct Submission
Submitted (108-JAN-2002) Institute of Zoology, Academia Sinica, 128
Academia Road, NanKang, Taipei 11529, Taiwan, ROC
Location/Qualifiers
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Detent: JP 2001275682-A 9 09-OCT-2001;
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PAT 08-OCT-2004

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geneseqn2003cs:\* geneseqn2003ds:\* geneseqn2004as:\*

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genesegn2004bs:\*genesegn2005s:\*genesegn2006s:\*

| Description                   | Aaf55098 DNA encod<br>Abi99041 Murine pc | Aax87813 HLA-DR2 a<br>bdw44282 DR2-19G f | Abi99027 IAS MBP 1 | Abi99033 MBP 90-10<br>Ado07566 Fusion or | Ado07578 Fusion pr | Ado07577 Fusion pr | Ad115694 Murine im<br>Aec20762 M-CSF spe | Aed19725 Anti-PrP | Adv26108 Mouse OKT | Adw71834 Murine OK | Aaq12637 Monoclona | Adq91058 Murine OK | Adt77690 Monoclona |
|-------------------------------|------------------------------------------|------------------------------------------|--------------------|------------------------------------------|--------------------|--------------------|------------------------------------------|-------------------|--------------------|--------------------|--------------------|--------------------|--------------------|
| SUMMARIES                     | AAF55098<br>ABI99041                     | AAX87813                                 | AB199027           | AB199033                                 | AD007578           | AD007577           | ADL15694<br>AEC20762                     | AED19725          | ADV26108           | ADW71834           | AAQ12637           | ADQ91058           | ADT77690           |
| DB                            | Ω4.α                                     | 4 6 5                                    | 4                  | 4 5                                      | 12                 | 12                 | 17                                       | 14                | 14                 | 14                 | 0                  | 12                 | 13                 |
| *<br>Query<br>Match Length DB | 1484                                     | 1446                                     | 2346               | 2343                                     | 1108               | 1108               | 990<br>1401                              | 1560              | 1569               | 1569               | 1570               | 1570               | 3973               |
| &<br>Query<br>Match           | 99.9                                     | . 4. 4.<br>                              | 48.9               | 48.7                                     | 47.2               | 47.2               | 47.2                                     | 47.2              | 47.2               | 47.2               | 47.2               | 47.2               | 47.2               |
| Score                         | 1484                                     | 957.6                                    | 726.2              | 723.2                                    | 701.2              | 701.2              | 700.6                                    | 700.6             | 700.6              | 700.6              | 700.6              | 700.6              | 200.6              |
| Fult<br>No.                   | 401                                      | J 44. R                                  | υ                  | r a                                      | 9                  | 10                 | 11                                       | 13                | 14                 | 15                 | 16                 | 17                 | 18                 |

Example 1; Page 31-33; 43pp; French.

| Aee21942 Single ch | Aaz35/04 Human gly | Ade85817 Murine in | Aat59350 1-153 del | Aat59349 1-153 c-m | Aaf30341 Bicistron | Aaf30316 Bicistron | Aeb12356 Fusion pr | Aed64236 mFc-hOGH- | Aef05391 Human mFc | Aeb12344 Immunoglo | Aed64224 Murine im | Aan91659 Chimeric | Aeb12360 Fusion pr | Aed64240 hTSH-mFC- | Aeb12362 Fusion pr | Aed64242 hOGH-mPC- | Aaq48037 Monoclona | Aaq54652 T84.12 He | Aav55416 Chimeric | Aav55415 Chimeric | Aat62850 Mouse sol | Ads31748 DNA encod |            | Aee21946 Single ch | Aaz51300 Murine im | Aaa50055 Mouse imm |
|--------------------|--------------------|--------------------|--------------------|--------------------|--------------------|--------------------|--------------------|--------------------|--------------------|--------------------|--------------------|-------------------|--------------------|--------------------|--------------------|--------------------|--------------------|--------------------|-------------------|-------------------|--------------------|--------------------|------------|--------------------|--------------------|--------------------|
| 4 AEE21942         | AAZ35/04           | 0 ADE85817         | AAT59350           | AAT59349           | AAF30341           | AAF30316           | 4 AEB12356         | 4 AED64236         | 5 AEF05391         | 4 AEB12344         | 4 AED64224         | AAN91659          | 4 AEB12360         | 4 AED64240         | 4 AEB12362         | 4 AED64242         | AAQ48037           | AAQ54652           | AAV55416          | AAV55415          | AAT62850           | 3 ADS31748         | 3 ADS92750 | 4 AEE21946         | AAZ51300           | AAA50055           |
| 1509 1             | 729 3              | 1140 10            | 1158 2             | 1188 2             | 6729 4             | 7528 4             | 1530 1             | 1530 1             | 699 1              | 708 1              | 708 1              | 1341 1            | 1413 1/            | 1413 14            | 1431 1,            | 1431 1             | 1581 2             | 1645 2             | 1131 2            | 1194 2            | 1275 2             | 1473 13            | 1473 1     | 1524 1,            | 699                | 699                |
| 47.2               | 47.2               |                    | ۲.                 | _                  |                    | _                  |                    | _                  |                    | _                  |                    |                   |                    | _                  | 47.1               | 47.1               | 47.1               | 47.1               | 47.0              | 47.0              | 47.0               | 47.0               | 47.0       | 47.0               | 47.0               | 47.0               |
| 700.4              | 700.2              | 700.2              | 200                | 700                | 700                | 700                | 699.4              | 699.4              | 669                | 669                | 669                | 669               | 669                | 669                | 669                | 669                | 669                | 669                | 698.4             | 698.4             | 698.4              | 698.4              | 698.4      | 697.8              | 697.4              | 697.4              |
| 19                 | 20                 | 21                 | 22                 | 23                 | 24                 | 25                 | 56                 | 27                 | 28                 | 53                 | 30                 | 31                | 32                 | 33                 | 34                 | 35                 | 36                 | 37                 | 38                | 33                | 40                 | 41                 | 42         | 43                 | 44                 | 45                 |

## ALIGNMENTS

New soluble recombinant protein, useful e.g. as immunostimulant, comprises dimeric major histocompatibility complex molecule fused to immunoglobulin Fc region. Recombinant protein; alpha chain; beta chain; MHC; immunoglobulin; major histocompatibility complex; Fc region; antigen; T lymphocyte; immunostimulant; vaccine; infection; tumour; ss. DNA encoding a fusion protein comprising an alpha chain of MHC. Location/Qualifiers 1..1482 /\*tag= a AAF55098 standard; DNA; 1484 BP. (CNRS ) CNRS CENT NAT RECH SCI. 29-JUL-1999; 99FR-00009862. 28-JUL-2000; 2000WO-FR002193 Glaichenhaus N, Malherbe L; 15-MAY-2001 (first entry) WPI; 2001-182944/18. P-PSDB; AAB67480. WO200109194-A1. 08-FEB-2001. Synthetic. AAF55098; 

The specification describes soluble recombinant proteins that comprise at least a dimer formed from the alpha and beta-chains of MHC (major histocompatibility complex) Class I and II mollecules in which at least one chain has, attached to its C-terminus, at least part of the Fc region of an immunoglobulin. The recombinant proteins, when linked to an antigenite peptide, are used to count and/or purify antigen-reactive T lymphocytes and to characterize their phenotype, e.g. in preclinical evaluation of vaccines. They are also used as immunostimulants, particularly for vaccine development (against infections and tumours), to count and determine phenotype of autoreactive T cells in subjects with, or at risk of developing, autoimmune diseases, e.g. for staging or evaluating treatments, and (to purify and/or enrich Ag-reactive T cells from cell cultures or patient samples, for use in subsequent curative or preventative cellular therapy. The present sequence encodes a recombinant protein of the invention, comprising an alpha chain of MHC molecules

Sequence 1484 BP; 414 A; 394 C; 362 G; 314 T; 0 U; 0 Other;

ö 120 240 240 300 300 360 420 420 480 480 540 900 720 540 9 9 720 780 TGCGGAGGTGAAGACGACATTGAGGCCGACGTAGGCTTCTATGGTACAACTGTTTAT 120 CAGTCTCCTGGAGACATTGGCCAGTACACACATGAATTTGATGGTGATGAGTTGTTCTAT 180 9 780 9 ATGCCGTGCAGCAGCAGAGCTCTGATTCTGGGGGTCCTCGCCCTGAACACCCTGGCCTCAGCCTC 61 TGCGGAGGTGAAGACGACATTGAGGCCGACCACGTAGGCTTCTATGGTACAACTGTTAT CAGTCTCCTGGAGACATTGGCCCAGTACACACATGAATTTGATGGTGATGATGTTCTAT GIGGACTIGGAIAAGAAAACTGICIGGAGGCTICCTGAGITTGGCCAAITGATACTC GAGACCAGCTTCCTCGTCAACCGTGACCATTCCTTCCACCAGCTGTCTTATCTCACCCTTC GGTGGAAGATCCACTACAGCTCCATCAGCTCAGCTCGAAAAAGAGCTCCAAGACCTCGGAG AAGGAAAATGCACAGCTGGAATGGGAGTTGCAAGCACTGGAAAAGGAACTGGCTCAGGCA AAGGAAAATGCACAGCTGGAATGGGAGTTGCAAGCACTGGAAAAAGGAACTGGCTCAGGCA 1 ATGCCGTGCAGCAGCACTCTGATTCTGGGGGTCCTCGCCCTGAACACCATGCTCAGCCTC Gregactregaraagaaaacrercregagecrrecreagrrresceaarrearacre ACTAAGAGGTCAAATTTCACCCCAGCTACCAATGAGGCTCCTCAAGGGACTGTGTTCCCC actrandadetenantricaceeecaderaceantendeecreerendeerererere AAGTCCCCTGTGCTGCTGGGTCAGCCCAACACCCTTATCTGCTTTTGTGACAACATCTTC GAGACCAGCTTCCTCGTCAACCGTGACCATTCCTTCCACAGAGCTGTCTTATCTCACCTTC ATCCCTTCTGATGATGACATTTATGACTGCAAGGTGGAGCACTGGGGCCTGGAGGAGCCG GTTCTGAAACATGGGAACCTGAGATTCCAGCCCCCATGTCAGAGCTGACAGAAACTGGA GGTGGAGGATCCACTACAGCTCCATCAGCTCCAGCTCGAAAAAGAGCTCCAGGCCCTGGAG Gaps ö ery Match 99.9%; Score 1484; DB 5; Length 1484; st Local Similarity 100.0%; Pred. No. 0; tches 1484; Conservative 0; Mismatches 0; Indels 0. 61 121 181 181 241 241 301 301 361 361 421 421 481 481 541 541 601 601 661 661 721

| \$ a                           | 781 (                                                         | 84<br>84                                                                                                                                                                                                                                                                                                                                                                                                      |                                                                             |
|--------------------------------|---------------------------------------------------------------|---------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|-----------------------------------------------------------------------------|
| & A                            | 841                                                           | AACCTCTTGGGTGGACCATCCGTCTTCATCTTCCCTCCAAAGATCAAGGATGTACTCATG 900<br>                                                                                                                                                                                                                                                                                                                                          |                                                                             |
| è 8                            | 901                                                           | 96                                                                                                                                                                                                                                                                                                                                                                                                            |                                                                             |
| දු දු                          | 961 (                                                         | 2 2                                                                                                                                                                                                                                                                                                                                                                                                           | 020                                                                         |
| දු ද                           | 1021                                                          | AGAGAGGATTACAACAGTACTCTCCGGGTGGTCAGTGCCCTCCCCCATCCAGCACCAGAC 106<br>                                                                                                                                                                                                                                                                                                                                          | 080                                                                         |
| දු පු                          | 1081                                                          | tggatgagtggcaaggagttcaaatgcaaggtcaacaacaaagacctcccagggccatc 114<br>                                                                                                                                                                                                                                                                                                                                           | 140                                                                         |
| දු දු                          | 1141 (                                                        | GAGAGAACCATCTCAAAACCCAAAGGGTCAGTAAGAGCTCCACAGGTATATGTCTTGCCT 120<br>                                                                                                                                                                                                                                                                                                                                          | 200                                                                         |
| 중 음                            | 1201                                                          | ccaccagaagaagagatgactaagaaacaggtcacttgacttgcattggtcacttg 126<br>                                                                                                                                                                                                                                                                                                                                              | 260                                                                         |
| දි දි                          | 1261                                                          | atgectgaagacatttacgtggagtggaccaacaacgaagaaacagaggtaaactacaag 13;<br>                                                                                                                                                                                                                                                                                                                                          | 320                                                                         |
| දි දි                          | 1321                                                          | arcactgaaccagtcctggactctgatggttcttacttcatgtacagcagctgagagtg 136<br>                                                                                                                                                                                                                                                                                                                                           | 380                                                                         |
| è 8                            | 1381 (                                                        | GAAAAGAAGAACTGGGTGGAAAGAATAGCTACTCCTGTTCAGTGGTCCACGAGGGTCTG 144<br>                                                                                                                                                                                                                                                                                                                                           | 4 4 0 4 0 4 0 0 4 4 0 0 4 4 0 0 4 4 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 |
| & g                            | 1441 (                                                        | CACAATCACCACACGACTAAGAGCTTCTCCCGGGCTAA 1484<br>                                                                                                                                                                                                                                                                                                                                                               |                                                                             |
| RESU<br>ABI9<br>XX<br>AC<br>XX | 2<br>BI99041<br>BI99041;                                      | tandard; cD                                                                                                                                                                                                                                                                                                                                                                                                   |                                                                             |
| EXE;                           | 25-FEB-200<br>Murine pCB                                      | 02 (first entry)<br>B223 coding sequence.                                                                                                                                                                                                                                                                                                                                                                     |                                                                             |
| X                              | Mouse; MH<br>single ch<br>antianaem<br>autoimmun<br>myastheni | Mouse; MHC; major histocompatibility complex; MHC class II; multimer; single chain; immunosuppressive; antidiabetic; autiinflammatory; entianemato; antirheumatoid, antiarthritic; neuroprotective; vaccine; autoimmune disease; insulin dependent diabetes; multiple sclerosis; myasthenia gravis; pernicious anaemia; autoimmune encephalomyelitis; rheumatoid arthritis; systemic lupus erythematosus; ss. |                                                                             |
| \$88X                          | B Sp.<br>nthetic                                              | ,                                                                                                                                                                                                                                                                                                                                                                                                             |                                                                             |
| Z X G                          | WO2001702                                                     | 45-A1.<br>01.                                                                                                                                                                                                                                                                                                                                                                                                 |                                                                             |

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22-MAR-2001; 2001WO-US009616.
22-MAR-2000; 2000US-0191274P.
15-MAY-2000; 2000US-0264003P.
23-JAN-2001; 2001US-0264003P.
(CORI-) CORIXA CORP.
Carter D, Zhu S, Arimilli S, Wang A; WPI; 2001-616371/71.
P-PSDB; ABB56471.
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Multimeric complex for treating autoimmune diseases, comprises first and second single chain MHC class II molecules, each comprising alphal and betal domain linked through amino acid linker and multimerization domain.

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Disclosure, Page 115; 147pp; English.

The invention relates to a multimeric complex comprising a first recombinant single chain major histocompatibility complex (MHC) class In molecule and a second recombinant single chain MHC class II molecule, each comprising an alphal domain and a betal domain linked through an amino acid linker and a multimerisation domain. The first and the second multimeric complex. The complex is useful for treating autoimmune diseases. It is useful for treating insulin dependent diabetes, multiple sclerosis, myasthenia gravis, pernicious anaemia, autoimmune encephalomyelitis (RAB), rheumatoid arthritis and systemic lupus molecule of the invention

Sequence 1676 BP; 438 A; 470 C; 407 G; 361 T; 0 U; 0 Other;

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127 CCTGGAGACATTGGCCAGTACACACATGATTTGATGGTGATGAGTTGTTCTATGTGGAC 186
151 CCTGGAGACATTGGCCAGTACACATGAATTTGATGGTGATGAGTGGTTCTATGTGGAC 210
  246
  306
   307 AGGICAAATTICACCCCAGCTACCAATGAGGCTCCTCAAGCGACTGTGTTCCCCAAGTCC 366
  331 AGGTCAAPATTCCACCCCAGCTACCAATGAGGCTCCTCAAGGGACTGTGTTCCCCAAGTCC 390
  367 CCTGTGCTGCTGGGTCAGCCCAACACCCTTATCTGCTGGACAACATCTTCCCCACCT 426
  510
   546
   570
   909
  211 ricgaraagaagacrarcregarecriccreacrirescearrisacaagcrireac 270
   187 TTGGATAAGAAGAAACTGTCTGGAGGCTTCCTGAGTTTGGCCAATTGATACTCTTTGAG
  247 CCCCAAGGTGGACTGCAAAACATAGCTGCAGAAAAACACAACTTGGGAATCTTGACTAAG
   487 AGCTTCCTCGTCAACCGTGACCATTCCTTCCACAAGCTGTCTTATCTCACCTTCATCCCT
   511 AGCTTCCTTGTCAACCGTGACCATTCCTTCCACAAGCTGTCTTATCTCACCTTCATCCCT
   547 TCTGATGATGACATTTATGACTGCAAGGTGGAGCACTGGGGCCTGGAGGAGCAGGTTCTG
Ouery Match 71.9%; Score 1067.4; DB 4; Length 1676; Sest Local Similarity 83.5%; Pred. No. 2.9e-277; Autches 1319; Conservative 0; Mismatches 96; Indels 164; Gaps
  427
  451
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| 571  | TCTGACGATGATATTTATGACTGCAAGGTGGAGCACTGGGGGCCTGGAGGAGCCGGTTCTG 630                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             |
|------|-------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| 607  | AAACACTGGGAACCTGAGATTCCAGCCCCCATGTCAGAGCTGACAGAAACTGGAGGTGGA 666                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              |
| 159  | GGAT                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          |
| 691  |                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               |
| 671  | 670                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           |
| 751  | GGATGCCTGGTCAAGGGTTATTTCCCTGAGCCAGTGACCTTGACCTGGAACTCTGGATCC 810                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              |
| 671  | CHARLOCAL CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF THE CONTROL OF |
| 110  | CIGICCERGIGGEGIGGERCANCACCIICCCARGCIGICCIGCAGICIGACCICCACACACCCICCACACACA                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     |
| 871  | AGCTCAGTGACTGTAACCTGGAGCACCTGGCCAGCCAGTCCATCACCTGCAATGTGG 928                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 |
| 747  | GTTGCAAGCACTGGAAAAGGAACTGGCTCAGGCAGCATCTGAGCCCAGAAGGCCCACAAT 806                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              |
| 807  | CCAAGCCCTGTCCTCCATGCAAATGCCCAGCACCTAACCTCTTGGGGGGGACCATCGTCTT 866                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             |
| 987  |                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               |
| 1047 | CATCTTCCCTCCAAAGATCAAGGATGTACTCATGATCTCCCTGAGCCCCATAGTCACATG 926                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              |
| 927  | TGTGGTGGTGGTTGTGAGCGAGATGACCCAGATGTCCAGATCAGCTGGTTTGTGAACAA 986                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               |
| 987  |                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               |
| 1047 | GGTGGTCAGTGCCTCCCCATCCAGCACCAGGACTGGATGAGTGGCAAGGAGTTCAAATG 1106<br>                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          |
| 1107 | CAAGGICAACAAAGACCICCCAGGGCCCATCGAGAACCATCTCAAAACCCAAAGG 1166<br>                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              |
| 1167 |                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               |
| 1227 | ACAGGICACTCTGACCTGCATGGTCACAGACTTCATGCCTGAAGACATTTACGTGGAGTG 1286                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             |
| 1287 | GACCAACAACGGGAAAACAGAGCTAAACTACAAGAACACTGAACCAGTCCTGGACTCTGA 1346<br>                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         |
| 1347 | TGGTTCTTACTTCATGTACAGCAAGCTGGAAAAGAAGAAGAACTGGGTGGAAGAAA 1406<br>                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             |
| 1407 | TAGCTACTCCTGTTCAGTGGTCCACGAGGGTCTGCACAATCACCACACGACTAAGAGCTT 1466                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             |
| 1467 | 467 CTCCCGGACTCCGGGTAAA 1485<br>                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              |

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  1118
  1070
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  1238
  290
   410
  959
  710
   770
  1058
   350
   476
   536
   296
  701
  191
  818
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   This nucleotide sequences codes for a bivalent DR2 fusion protein obtained by fusion of the Pc portion of 19G2a to the 3' end of a DR-alpha PCS cDNA construct (see AAV16866). The Pc portion was amplified by RT-PCR from mouse hybridoma 1243. The PCR product was then fused in frame with the DR alpha-POS construct by overlapping PCR. The DR2-19G fusion was expressed in the Drosophila Schneider cell system. The invention relates to new soluble monovalent and multivalent Class II MHC fusion proteins comprising a MHC class II binding domain and a dimerisation domain or an immunoglobulin region that can be used for the treatment of allergic and autoimmune diseases (e.g. multiple sclerosis), for tolerising a subject to foreign tissue before or after organ or tissue transplantation, or for vaccination against pathogens. (Updated on 17-OCT-2003 to standardise OS field)
   116 TITATCAGICICCIGGAGACATIGGCCAGIACACACAIGAATTIGAIGGIGAIGAIGT 175
   176 TCTATGTGGACTTGGATAAGAAAGTGTCTGGAGGCTTCCTGAGTTTGGCCAATTGA 235
  110 rccardricianaridecanadandencedecriciacedecricandantricandaring 169
   Major histocompatibility complex class II; MHC class II; human; mouse; fusion protein; HLA-PRZ; DRA*0101; binding domain; Fos; dimerisation domain; IGG; allergy; autoimmune disease; vaccine; multiple sclerosis; therapy; ss.
  CCAGCTTTGAGGCTCAAGGTGCATTGGCCAACATAGCTGTGGAAAAGCCCAACTTGGAAA
  New Class II MHC fusion proteins - comprising a MHC Class II binding domain and a dimerisation domain or an immunoglobulin region used for modulating immune responses.
   236 TACTCTTTGAGCCCCAAGGTGGACTGCAAAACATAGCTGCAGAAAAACACACAACTTGGGAA
  Gaps
   Indels 18;
   Score 957.6; DB 2; Length 1446;
Pred. No. 1.2e-247;
0; Mismatches 234; Indels 18;
  Sequence 1446 BP; 414 A; 375 C; 356 G; 301 T; 0 U; 0 Other;
   Ë
   Example; Page 49; 76pp; English
   Wucherpfennig KW, Strominger
   AAT99707 standard; cDNA; 1446
   64.5%;
  96US-0024077P
  97WO-US014503
   (revised)
(first entry)
  DR2-IgG fusion construct.
  tches 1136; Conservative
   (HARD ) HARVARD COLLEGE
   WPI; 1998-159459/14.
  Local Similarity
  Homo sapiens.
Mus musculus.
Chimeric.
  15-AUG-1997;
  16-AUG-1996;
  WO9806749-A2
   17-OCT-2003
17-AUG-1998
  19-FEB-1998,
   AAT99707;
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CTCCACAGGTATATGTCTTGCCTCCACAGAAGAAGAGATGACTAAGAAACAGGTCACTC 1237
  CCCTCCCCATCCAGCACCAGGACTGGATGAGTGCCAAGGAGTTCAAATGCAAGGTCAACA 1117
  ACAAAGACCTCCCAGCGCCCATCGAGAGAACCATCTCAAAAACCCAAAGGGTCAGTAAGAG 1177
   415
  535
  595
   997
   949
  469
   529
  655
   589
   700
   649
   760
  709
  817
  769
  877
   829
  937
   889
   TCCCCAAGTCCCCTGTGCTGCTCAGCCCCAACACCCTTATCTGCTTTTGTGGACAACA
  rercacidadescrircirciscociassandiciacerrirciscandricacinitric
  cerrecrecercaacreageacerrraceacrecageaceageacresescrresars
   ATGTGAGCGAGGATGACCCAGATGCCAGATCAGCTTGTTGTGAACAACGTGGAAGTAC
  ACACAGCTCAGACACAAACCCATAGAGAGATTACAACAGTACTCTCCGGGTGGTCAGTG
  CCTTCATCCCTTCTGATGATGACATTTATGACTGCAAGGTGGAGCACTGGGGCCTGGAGG
   530 AGCCTCTTCTCAAGCACTGGGAGTTTGATGCTCCAAAGCCCTCTCCCAGAGACTACAGAGG
   CTCCATGCAAATGCCCAGCACCTAACCTCTTGGGTGGACCATCCGTCTTCATCTTCCCTC
  CAAAGATCAAGGATGTACTCATGATCTCCCTGAGCCCCCATAGTCACATGTGTGGTGGTGG
   TGACCTGCATGGTCACAGACTTCATGCCTGAAGACATTTACGTGGAGTGGACCAACAACG
  AGCCGGTTCTGAAACACTGGGAACCTGAGATTCCAGCCCCCATGTCAGAGCTGACAGAAA
   CTGGAGGTGGAGGATCCACT------ACAGCTCCATCAGCTCGAAA
   590 rcéacecacicices de restrante de la reconsola de la reconsta del reconsta de la reconsta de la reconsta del reconsta de la reconsta del reconsta de la reconsta de la reconsta de la reconsta de la reconsta del reconsta de la reconsta del reconsta de la reconsta de la reconsta de la reconsta de la reconsta del reconsta de la reconsta de la reconsta de la reconsta
  AAAAGGAACTG---GCTCAGGCAGCATCTGAGCCCCAGAGGGCCCCACAATCAAGCCCTGTC
   CCCTCCCCATCCAGCACCAGGACTGGATGAGTGGCAAGGAGTTCAAATGCAAGGTCAACA
TCTTGACTAAGAGGTCAAATTTCACCCCAGCTACCAATGAGGCTCCTCAAGCGACTGTGT
   AAGAGCTCCAGGCCCTGGAGAAGGAAATGCACAGCTGGAATGGGAGTTGCAAGCACTGG
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1369
   Major histocompatability complex Class II; MHC; binding domain; HIA-DR2; leucine zipper; Fos; IgG; Fc; immunoglobulin; antibody; fusion protein; multiple sclerosis; rheumatoid arthritis; graft rejection; allergy; autoimmune disease; pemphigus vulgaris; systemic lupus erythematosus; T cell; diagnosis; therapy; adoptive immunotherapy; ss.
New HMC Class II binding domain fusion proteins and conjugates - used for, e.g. treating allergic and autoimmune diseases or detecting, isolating, activating or killing specific T cells.
  /note= "alpha-mating factor secretion signal"
16. 1437
/*tag= c
/product= "DR2-Fos-Fc"
  HLA-DR2 alpha-Fos-IgG fusion construct.
  Location/Qualifiers
1. .1440
/*tag= a
   BP
  Wucherpfennig KW, Strominger
   AAX87813 standard; DNA; 1446
   99WO-US003603
   98US-0075351P
  (first entry)
   Д
  cerevisiae
  1. .15
/*tag=
   (HARD ) HARVARD COLLEGE
   CGGCTAAA 1485
   1430 CGGGTAAA 1437
  WPI; 1999-527481/44.
   P-PSDB, AAY31654.
   Homo sapiens.
Saccharomyces c
Synthetic.
Chimeric.
  WO9942597-A1
   19-FEB-1999;
   19-FEB-1998;
  09-NOV-1999
  sig_peptide
   mat_peptide
   1478
   AAX87813;
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This nucleotide sequence codes for a divalent HIA-DR2 MHC binding domain fusion protein (see AAY31654) comprising an alpha-mating factor secretion signal, the extracellular domain of the HIA-DR2 alpha chain (residues 1-191 of DRA\*0101), a 7-amino acid linker, the 40-amino acid leucine zipper dimerization domain of Fos, and the Fc portion of IgG2a. The DR-alpha-Fc chain corresponds to an antibody heavy chain. The invention provides new monovalent, multivalent and multimeric MHC class II binding domain fusion proteins and conjugates comprising at least a binding domain of an MHC

Example 7, Page 100-102; 113pp; English.

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criass II alpha or beta chain and a dimerization domain, especially a Fos or Jun leucine zipper domain. The MHC fusion proteins and conjugates can be used if for detecting and isolating T cells having a defined MHC/peptide complex specificity (claimed); to confer to a subject adoptive immunity to a defined MHC/peptide complex (claimed) is to stimulate or activate T cells reactive to a defined MHC/peptide complex (claimed); for selective killing of T cells reactive to a defined MHC/peptide complex (claimed); to treat claiming of T cells reactive to a defined MHC/peptide complex (claimed); to treat allergic and autoimmune diseases, e.g. multiple sclerosis, rheumatoid arthritis, pemphigus vulgaris, and systemic lupus erythematosus; and to prevent organ or tissue transplant rejection. The DR2-IgG design was chosen to increase the affinity for the T cell receptor by increasing valency, and to attach an effector domain, the Rc region of IgG2a.

Complement fixation may result in the lysis of target T cells following binding of DR2-IgG molecules can any therefore be useful for the selective depletion of autoaggressive T
  235
  355
  230 rcarcacaaaccerrccaacraracrcccarcaccarcacrccrccacaccracrerer 289
   415
  409
  469
  595
   655
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   649
   760
  709
  169
  817
  CTCCATGCAAATGCCCAGCACCTAACCTCTTGGGTGGACCATCCGTCTTCATCTTCCCTC 877
   AGCCTCTTCTCAAGCACTAGGGAGTTTGATGCTCCAAGGCCTCTCCCAGAGACTACAGAGG
  AGTTCATCCTGGCCGCCCATGCAGCATCTGAGCCCAGAGGGCCCACAATCAAGCCCTGTC
  TCTATGTGGACTTGGATAAGAAAACTGTCTGGAGGCTTCCTGAGTTTGGCAATTGA
  110 rccargricanaris de la canada de la compani
  ccaectringagecreaagenecarnegecaacaracarecreacaaaageccaactregaaa
   TCCCCAAGTCCCCTGTGCTGCTCAGCCCAACACCCTTATCTGCTTTGTGGACAACA
  rcaccccaccagregical architectus de la consecue de la
  TITATGAGACCAGCITCCTCGTCAACCGTGACCATTCCTTCCACAAGCTGTCTTATCTCA
   cerricarecerricardardacarrirardaciaedadegenegagedecregage
   CCTTCCTGCCCTCAACTGAGGACGTTTACGACTGCAGGTGGAGGACTGGGGGTGGACTGGGTG
   AGCCGGTTCTGAAACACTGGGAACCTGAGATTCCAGCCCCCATGTCAGAGCTGACAGAAA
   CTGGAGGTGGAGGATCCACT------ACAGCTCCATCAGCTCGAAA
   590 redacedaciosececeerrraacrearacaceceaseceaseacacacicaac
   aceaeaacrenecerie ca de construir esta de construir de co
  236 TACTCTTTGAGCCCCAAGGTGGACTGCAAAACATAGCTGCAGAAAAACACAACTTGGGAA
   TCTTGACTAAGAGGTCAAATTTCACCCCAGCTACCAATGAGGCTCCTCAAGCGACTGTGT
  AAGAGCTCCAGGCCCTGGAQAAAAGGAAAATGCACAGGAATGGGAATTGGAGTTGCAAGCACTGG
  AAAAGGAACTG - - - GCTCAGGCAGCATCTGAGCCCAGAGGGCCCACAATCAAGCCCTGTC
   116 TTTATCAGTCTCCTGGAGACATTGGCCAGTACACACATGAATTTGATGGTGATGAGTTGT
  50 rcrarcraarccraarcaaccaarcaaccaarrrargraraargargargarrarrr
   Gaps
   18;
  Score 957.6; DB 2; Length 1446; Pred. No. 1.2e-247; 0; Mismatches 234; Indels 18;
  Sequence 1446 BP; 414 A; 375 C; 356 G; 301 T; 0 U; 0 Other;
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Best Local Similarity 81.8%;
Matches 1136; Conservative (
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   290
  416
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   530
  959
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1070 1178 1130 1238 1190 1298 1250

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110 recardidearardecaaadaadaadaadacerececerreaadaarrideacearire 169
   The present invention relates to the class II major histocompatibility complex (MHC) fusion protein having MHC class II binding domain of MHC class II alpha chain and a dimerization domain. The invention is useful in adoptive immunotherapy and tolerizing against foreign tissue. The invention is also useful for treating autoimmune diseases such as pemphigus vulgaris, rheumatoid arthritis, multiple sclerosis and systemillupus erythematosus. The present sequence is the DR2-IGG fusion protein
   Novel class II major histocompatibility complex (MHC) fusion protein having MHC class II binding domain of MHC class II alpha chain, and dimerization domain, useful for treating pemphigus vulgaris, rheumatoid
  rcrarcraarccrarcadecdadrrrargrridacriridarderedadarrr
  176 TCTATGTGGACTTGGATAAGAAAAACTGTCTGGAGGCTTCCTGAGTTTGGCCAATTGA
   236 TACTCTTTGAGCCCCAAGGTGGACTGCAAAACATAGCTGCAGAAAAACACACTTGGGAA
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589. .609
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   Leucine zipper domain"
   64.5%; Score 957.6; DB 14;
llarity 81.8%; Pred. No. 1.2e-247;
Conservative 0; Mismatches 234;
  secretory
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  Example; SEQ ID NO 11; 55pp; English.
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   ij
   ö
  end
  96US-0024077P.
97WO-US014503.
98US-0075351P.
99US-00248964.
   Wucherpfennig KW, Strominger
  2004US-00895543
No
No
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   ×
           1. .15
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  WPI; 2005-089945/10.
   Best Local Similarity
Matches 1136; Conserv
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  US2005003431-A1
   16-AUG-1996;
15-AUG-1997;
19-FEB-1998;
  21-JUL-2004;
                misc_feature
   misc_feature
  misc_feature
   misc_feature
   misc feature
   12-FEB-1999;
   DNA
  06-JAN-2005
  arthritie
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  230
   encoding
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   1237
  1297
   1429
  1057
   1009
   1357
  1309
   889
  997
   adoptive immunotherapy; dermatological; immunosuppressive; antirheumatic; antiatrhitic; neuroprotective; antiinflammatory; autoimmune diseases; pemphigus vulgaris; rheumatoid arthitis; multiple aclerosis; systemic lupus erythematosus; immune disorder; DR2-1gg protein; gene; ds. systemic lupus
   CCCTCCCCATCCAGCACCAGGACTGGATGAGTGGCAAGGAGTTCAAATGCAAGGTCAACA
  ACAAAGACCTCCCAGCGCCCATCGAGAGAACCATCTCAAAACCCAAAGGGTCAGTAAGAG
  Arcreacceacearceacarcreatercacarcacrecrrecrreaceaccaccreaacrac
   ACACAGCTCAGACACAAACCCCATAGAGAGGATTACAACAGTACTCTCCGGGTGGTCAGTG
   ACACAGCTCAGACACAAACCCATAGAGAGGATTACAACAGTACTCTCCGGGTGGTCAGTG
   CCCTCCCCATCCAGCACCAGGACTGGATGAGTGGCAAGGAGTTCAAATGCAAGGTCAACA
  ACAAAGACCTCCCAGCGCCCATCGAGAACCATCTCAAAACCCAAAGGGTCAGTAAGAG
   CTCCACAGGTATATGTCTTGCCTCCACCAGAAGAGAGATGACTAAGAAACAGGTCACTC
   TGACCTGCATGGTCACAGACTTCATGCCTGAAGACATTTACGTGGAGTGGACCAACAACG
   TGACCTGCATGGTCACACACACTTCATGCCTGAAGACATTACGTGGAGTGGACCAACAACG
   GGAAAACAGAGCTAAAACTACAAGAACACTGAACCAGTCCTGGACTCTTGATGGTTCTTACT
  GGAAAACAGAGCTAAAACTACAAGAACACTGAACCAGTCCTGGACTCTGATGGTTCTTACT
   Major histocompatibility complex; fusion protein; immunoconjugate;
  1. .1440
/trag= b
/product= "DR2-IgG fusion protein"
/partial
   DR2-IgG fusion protein encoding DNA
  Location/Qualifiers
  B
  ADW44282 standard; DNA; 1446
   (first entry)
  CGGGTAAA 1437
   CGGGTAAA 1485
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1310 1418 1370 1478 1430

1358

ä 175 109 235

18; 'Gaps

IndelB

295 229 355

sapiens Chimeric. Unidentified

Ношо

Key

24-MAR-2005

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GTTCAGTGGTCCACGAGGGTCTGCACAATCACCACGACGACTAAGAGCTTCTCCCCGGACTC
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  22-MAR-2000; 2000US-0191274P.
15-MAY-2000; 2000US-0204249P.
23-JAN-2001; 2001US-0264003P.
   22-MAR-2001; 2001WO-US009616
  Query Match
Best Local Similarity 70.5%;
Matches 1095; Conservative
  Zhu S, Arimilli
  (first entry)
   molecule of the invention
   1485
  1437
  WPI; 2001-616371/71.
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   CGGGTAAA
  CCCCTAAA
  P-PSDB; ABB56457
   WO200170245-A1
  25-FEB-2002
  27-SEP-2001
  Carter D,
  Synthetic.
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  ABI99027;
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   1249
  1057
  1117
  1069
  1117
  1129
   1237
  1189
   1297
  1357
   1417
   949
                             TCACGAACAGCCCTGTGGAACTGAGAGGCCCAACGTCCTCATCTGTTTCATAGACAAGT 349
  409
   469
   595
  529
   655
   700
  649
  760
   709
   817
   769
   877
   829
   937
  889
  997
              TCCCCAAGTCCCCTGTGCTGGGTCAGCCCAACACCCTTATCTGCTTTGTGGACAACA
  TCACCCCACCAGTGGTCAATGTCACGTGGCTTCGAAATGGAAAACCTGTCACCACAGGAG
   TITATGAGACCAGCTICCTCGTCAACCGTGACCATTCCTTCCACAAGCTGTCTTATCTCA
   TGTCAGAGACAGTCTTCCTGCCCAGGGAAGACCACCTTTTCCGCAAGTTCCACTATCTCC
   CCTTCATCCCTTCTGATGATGACATTTATGACTGCAAGGTGGAGCACTGGGGCCTGGAGG
  AGCCGCTTCTCAAACACTGGGAACCTGAGATTCCAGCCCCCATGTCAGAGCTGACAGAAA
   CTGGAGGTGGAGGATCCACT------ACAGCTCCATCAGCTCGAAA
   TCGACGGAGGTGGCGCGGTTTAACTGATACACTCCAAGCGGAGACAGATCAACTTGAAG
   650 ACGAGAAGTCTGCGTTGCAGACCGAGATTGCCAATCTACTGAAAGAGAAGGAAAAAACTGG
  Agricarccroscoccocarscascarcrosascocasascoccacaarcaascocroro
  CAAAGATCAAGGATGTACTCATGATCTCCCTGAGCCCCATAGTCACATGTGTGGTGGTGG
  ATGTGAGGGATGACCCAGATGTCCAGATCAGCTGGTTTGTGAACAACGTGGAAGTAC
  CCTCCCCATCCAGCACCAGGACTGGATGAGTGGCAAGGAGTTCAAATGCAAGGTCAACA
   ACAAAAACCTCCCAGCGCCCATCGAGAGAACCATCTCAAAAACCCAAAAGGGTCAGTAAGAG
  ACAAAGACCTCCCAGCGCCCATCGAGAGCCATCTCAAAACCCAAAGGGTCAGTAAGAG
   TGACCTGCATGGTCACAGACTTCATGCCTGAAGACATTTACGTGGAGTGGACCAACACG
   GGAAAACAGAGCTAAAACTACAAGAACACTGAACCAGTCCTGGACTCTTGATGGTTCTTACT
   AGCCTCTTCTCAAGCACTGGGAGTTTGATGCTCCAAGCCCCTCTCCCAGAGACTACAGAGG
  AAGAGCTCCAGGCCCTGGAGAAAGGAAAATGCACAGCTGGAATGGGAGTTGCAAGCACTGG
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  CAAAGATCAAGGATGTACTCATGATCTCCCTGAGCCCCCATAGTCACATGTGTGGTGGTGGTGG
   ACACAGCTCAGACACAAACCCATAGAGAATTACAACAGTACTCTCCGGGTGGTCAGTG
  CCCTCCCCATCCAGCACCAGGACTGGATGAGTGGCAAGGAGTTCAAATGCAAGGTCAACA
  CTCCACAGGTATATGTCTTGCCTCCACAGAAGAAGAAGAGATGACTAAGAAACAGGTCACTC
  crecacadenararererrecerceacadaadaadaacadaaaaaacaderee
   TGACCTGCATGGTCACAGACTTCATGCCTGAAGACATTTACGTGGAGTGGACCAACAACG
   GGAAAACAGAGCTAAACTACAAGAACACTGAACCAGTCCTGGACTCTGATGGTTCTTACT
   CTCCATGCAAATGCCCAGCACCTAACCTCTTGGGTGGACCATCCGTCTTCATCTTCCCTC
  AAAAGGAACTG---GCTCAGGCAGCATCTGAGCCCCAGAGGGCCCACAATCAAGCCCTGTC
   290
  710
  818
  770
  878
  830
   950
   1058
  1010
   1118
   1070
   1178
   1130
   1238
   1190
  1358
  1310
                                     290
   350
   476
   410
   536
   296
   530
   959
  701
  938
   966
   416
  191
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4
  The invention relates to a multimeric complex comprising a first recombinant single chain major histocompatibility complex (MHC) class II molecule and a second recombinant single chain MHC class II molecule, each comprising an alphal domain and a betal domain linked through an amino acid linker and a multimerisation domain. The first and the second molecule are linked through the multimerisation domain. The first and the second multimeric complex, The complex is useful for treating autodimmune diseases. It is useful for treating insulin dependent diabetes, multiple solerosis, myasthenia gravis, pernicious ansemia, autodimmune encephalomyelitis (BAE), rheumatoid arthritis and systemic lupus erythematogus. The present sequence encodes a single chain MHC class II
  Multimeric complex for treating autoimmune diseases, comprises first and second single chain MHC class II molecules, each comprising alphal and betal domain linked through amino acid linker and multimerization domain.
   Mouse; MHC; major histocompatibility complex; MHC class II; multimer; single chain; immunosuppressive; antidiabetic; antiinflammatory; antiansemic; antirheumatoid; antiarthritic; neuroprotective; vaccine; autoimmune disease; insulin dependent diabetes; multiple sclerosis; myasthenia gravis; pernicious anaemia; autoimmune encephalomyelitis; rheumatoid arthritis; systemic lupus erythematosus; ss.
  Score 726.2; DB 4; Length 2346;
Pred. No. 4.1e-185;
0; Mismatches 323; Indels 135;
  Sequence 2346 BP; 560 A; 663 C; 646 G; 477 T; 0 U; 0 Other;
IAS MBP 1-14 CH1. CH2. CH3 coding sequence.
  Ä
   Disclosure; Page 91-92; 147pp; English
  Wang
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TGAGAGTGGAAAAAAAAGAACTGGGTGGAAAGAAATAGCTACTCCTGTTCAGTGGTCCACG 1432
   Multimeric complex for treating autoimmune diseases, comprises first and second single chain MHC class II molecules, each comprising alphal and betal domain linked through amino acid linker and multimerization domain.
                               ACCAGGACTGGATGAGTGGCAAGGAGTTCAAATGCAAGGTCAACAACAAAGACCTCCCAG
   cadactrictricchdaddacatracricricdadriodcagrogaardddcagccadccigada
   acracaagaacacrcagcccarcarggacacagarggcrcrracrrcgrcracaggaagc
  Mouse, MHC, major histocompatibility complex; MHC class II; multimer; single chain; immunosuppressive; antidiabetic; antinflammatory; antianaemic; antirheumatoid; antiarthritic; neuroprotective; vaccine; autoimmune disease; insulin dependent diabetes; multiple sclerosis; mayaethenia gravis; pernicious anaemia; autoimmune encephalomyelitis; rheumatoid arthritis; systemic lupus erythematosus; ss.
  TCTTGCCTCCACCAGAAGAAGAGATGACTAAGAAACAGGTCACTCTGACCTGCATGGTCA
   CAGACTTCATGCCTGAAGACATTTACGTGGAGTGGACCAACAACGGGAAAACAGAGCTAA
   ACTACAAGAACACTGAACCAGTCCTGGACTCTGATGGTTCTTACTTCATGTACAGCAAGC
   AGGGTCTGCACAATCACCACACGACTAAGAGCTTCTCCCGGACTCCGGGTAAA
   CGCCCATCGAGAGAACCATCTCAAAACCCAAAGGGTCAGTAAGAGCTCCACA
   90-101 CHI.H.CH2.CH3 coding sequence
   ä
  Wang
   Disclosure; Page 96; 147pp; English
  BP.
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  ABI99033 standard; cDNA; 2343
   22-MAR-2000; 2000US-0191274P.
15-MAY-2000; 2000US-0204249P.
23-JAN-2001; 2001US-0264003P.
   22-MAR-2001; 2001WO-US009616.
  Arimilli
  (first entry)
   WPI; 2001-616371/71.
  (CORI-) CORIXA CORP.
  Zhu S,
  P-PSDB; ABB56463
  WO200170245-A1.
  25-FEB-2002
   27-SEP-2001
   Mus sp.
Synthetic.
   2039
  2159
   'n
   1919
   2099
   2219
   1433
   2279
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   1253
  1313
   1373
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   1798
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  1918
   1138
  1318
  1498
  1018
   1078
  1198
   1258
   CAGTGACTGTCCCCTCCAGCACCTGGCCCAGCGAGCGTCACCTGCAACGTTGCCCACC 1618
   367
   427
  487
  547
   607
  618
   713
  735
  892
   952
  GCTTCCTTGTCAACCGTGACCATTCCTTCCACAAGCTGTCTTATCTCACCCTTCATCCCTT
  CTGACGATGATATTTATGACTGCAAGGTGGAGCACTGGGGCCTGGAGGGGGCGTTCTGA
   crercrarceacresciceresarcrecreceaaacraceareacrecressar
  CCTGGAGAAGGAAAAT-------GCACAG
   GTGAAGACGACATTGAGGCCGACCACGTAGGCGTCTATGGTACAACTGTATATCAGTCTC
   CTGGAGACATTGGCCAGTACACACATGAATTTGATGGTGATGAGTGGTTCTATGTGGACT
   TGGATAAGAAGAAAACTGTCTGGAGGCTTTCCTGAGTTTGGCCAATTGATACTCTTTGAGC
  regaraagaagacrarcregarecriccreacrirescearreacaaccrireace
  CCCAAGGTGGACTGCAAAACATAGCTGCAGAAAAACACAACTTGGGAATCTTGACTAAGA
   GGTCAAATTCCACCCCAGCTACCAATGAGGCTCCTCAAGCGACTGTGTTCCCCCAAGTCCC
   CTGTGCTGCTGGGTCAGCCCAACACCCTTATCTGCTTTGTGGACAACATCTTCCCACCTG
  TGATCAACATCACATGGCTCAGAAATAGCAAGTCAGTCACAGACGGCGTTTATGAGACCA
  TGATCAACATCACATGGCTCAGAAATAGTAAGTCAGTCACAGACGGCGTTTATGAGACCA
   GCTTCCTCGTCAACCGTGACCATTCCTTCCACAAGCTGTCTTATCTCACCTTCATCCCTT
   CTGATGATGACATTTATGACTGCAAGGTGGAGCACTGGGGGCCTGGAGGAGCCGGTTCTGA

    ACTGGAGGTGGAGGATCCACTACAGCTCCATCAGCTCCAAAAAAAGGCTCCAGGC

  1439 GCCTGGTCAAGGGCTATTTCCCTGAGCCAGTGACAGTGACCTGGAACTCTGGATCCCTGT
  ccadedegreracacerreceagererecreerereacereracacrereaddager
   CTGGAATGGGAGTTGCAAGCACTGGAAAAGGAACTGGCTCAGGCAGCATCTGAGCCCAGA
  cesceaecaeceaegresaeaagaresaeaaarrereceaegearreresaecerr
  ---CACCTAACCTCTTGGGTGGACCATCCGTCTTCATCTTCCCTCCAAAGATCAAGGATG
  gcanárgiacagicccagaagnárcaicricircírcárcircécccaaagcccaaggaig
   893 TACTCATGATCTCCCTGAGCCCCCATAGTCACATGTGTGGTGGTGGATGTGAGCGATG
   TGCTCACCATTACTCTGACTCCTAAGGTCACGTGTTGTGGTAGACATCAGCAAGGATG
  AAACCCATAGAGGATTACAACAGTACTCTCCGGGTGGTCAGTGCCCTCCCCATCCAGC
  796 GGCCCACAATCAAGCCCTGTCCTCCATGCAAATGCCCAG-----
   CCTGAGATTCCAGCCCCCATGTCAGAGCTGACAGAA----
  AACACTGGG------
  188
  1079
  608
   1319
  655
   1499
   1619
   836
   1739
  1013
   1019
   1139
   1199
  1259
   1379
   1679
 779
                               128
  899
  248
   308
  428
   488
  548
   619
   736
  953
```

The invention relates to a multimeric complex comprising a first recombinant single chain major histocompatibility complex (MHC) class II molecule and a second recombinant single chain MHC class II molecule, each comprising an alphal domain and a betal domain linked through an amino acid linker and a multimerisation domain. The first and the second molecule are linked through the multimerisation domain to form a multimeric complex. The multimeric complex, the complex is useful for treating autoimmune diseases. It is useful for treating insulin dependent diabetes, multiple sclerosis, myasthenia gravis, pernicious anaemia, autoimmune encephalomyelitis (RAB), rheumatoid arthritis and systemic lupus erythematosus. The present sequence encodes a single chain MHC class II molecule of the invention

Sequence 2343 BP; 562 A; 665 C; 635 G; 481 T; 0 U; 0 Other;

953 CCCAAGGTGGACAAACATAGCTACAGGAAAATACACCTTGGGAATCTTGACTAAGA 1012 1133 TGATCAACATCACATGGCTCAGAAATAGTAAGTCACACAGACGGCGTTTATGAGACCA 1192 1193 GCTTCCTTGTCAACCGTGACCATTCCTTCCACCAGCTGTCTTATCTCACCTTCATCCCTT 1252 1253 CTGACGATGATATTATGACTGCAAGGTGGAGCACTGGGGCCTGGAGGAGCCGGTTCTGA 1312 1013 GGTCAAATTCCACCCCAGCTACCAATGAGGCTCCTCAAGGGACTGTGTTCCCCAAGTCCC 1072 1073 crigracorcaccica acaccica referencia de contra con contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra contra co 1313 AACACTGGGCGAGGGGGGGGGGGGGGGGAGGAAGCTTAGCCAAAACGACACCCC 1372 1433 GATGCCTGGTCAAGGGCTATTTCCCTGAGCCAGTGACAGTGACCTGGAACTCTGGATCCC 1492 1493 TGTCCAGCGGTGTGCACACCTTCCCAGCTGTCCTGCAGTCTGACCTCTACACTCTGAGCA 1552 832 CCCAAGGTGGAAAACATAGCTGCAGAAAACACACAACTTGGGAATCTTGACTAAGA 307 607 710 68 GTGAAGACGACATTGAGGCCGACCACGTAGGCTTCTATGGTACAACTGTTTATCAGTCTC 127 892 GGTCAAATTTCACCCCAGCTACCAATGAGGCTCCTCAAGCGACTGTGTTCCCCAAGTCCC 367 427 487 547 616 732 128 CTGGAGACATTGGCCAGTACACACATGAATTTGATGGTGATGAGTTGTTCTATGTGGACT 187 188 TGGATAAGAAGAAAACTGTCTGAGCTTCCTGAGTTTGGCCAATTGATACTCTTTGAGC 247 792 655 ----ACTGGAGGTGGAGGATCCACTACAGCTCCATCAGCTCGGGAAAAAGAGCTCCA 733 CAGCTGGAATGGGAGTTGCAAGCACTGGAAAAGGAACTGGCTCAGGCAGCATCTGAGCCC CTGTGCTGCTGGGTCAGCCCAACACCCTTATCTGCTTTGTGGACAACATCTTCCCACCTG TGATCAACATCACATGGCTCAGAAATAGCAAGTCAGTCACAGACGGCGTTTATGAGACCA GCTTCCTCGTCAACCGTGACCATTCCTTCCACAAGCTGTCTTATCTCACCTTCATCCCTT 548 CTGATGATGACATTTATGACTGCAAGGTGGAGCACTGGGGCCTGGAGGAGCCGGTTCTGA Indels 138; Gaps Length 2343; 0; Mismatches 323; DB 4; 617 - AACCTGAGATTCCAGCCCCCATGTCAGAGCTGACAGAA 48.7%; Score 723.2; DB 4; 70.4%; Pred. No. 2.6e-184; 711 GGCCCTGGAGGAAGGAAAT---est Local Similarity 70.4 Vatches 1095; Conservative 608 AACACTGGG---248 368 uery Match 308 428 488

| CCTGCAACGTTGCCC 1613                            | 835                                           | ATTGTGGTTGTAAGC 1673 | CTCCAAAGATCAAGG 889                                     | CCCCAAAGCCCAAGG 1733                           | σ,                                                                 | TAGACATCAGCAAGG 1/9:                              |                                                                                                   | Grecerecearee 1069                           | <br> GTGAACTTCCCATCA 1912                     | ACAACAAAGACCTCC 1129<br>                                   | -  | 203  | CTCTGACCTGCATGG 1249                                           | GTCTGACCTGCATGA 2092                      | ACGGGAAACAGAGC 1309                                         | Areeckeckecke 2152                                 | 136                                                        | actrosrcracasca 2213                           | CCTGTTCAGTGGTCC 1429                                         | 1485 | 232      |                                |         |                    |                                          | <pre>antidiabetic;<br/>antiinflammatory;<br/>; gene.</pre>                                                                                                                   |                      |                                                                               |
|-------------------------------------------------|-----------------------------------------------|----------------------|---------------------------------------------------------|------------------------------------------------|--------------------------------------------------------------------|---------------------------------------------------|---------------------------------------------------------------------------------------------------|----------------------------------------------|-----------------------------------------------|------------------------------------------------------------|----|------|----------------------------------------------------------------|-------------------------------------------|-------------------------------------------------------------|----------------------------------------------------|------------------------------------------------------------|------------------------------------------------|--------------------------------------------------------------|------|----------|--------------------------------|---------|--------------------|------------------------------------------|------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|----------------------|-------------------------------------------------------------------------------|
| 3 GCTCAGTGACTGTCCCCTCCAGCACCTGGCCCAGCGAGACCGTCA | 3 AGAGGCCCACAATCAAGCCCTGTCCTCCATGCAAATGCCCAG- |                      | 6CACCTAACCTCTTGGGTGGACCATCCGTCTTCATCTTCCCTCCAAAGATCAAGG | 3 CTTGCATATGTACAGTCCCAGAAGTATCATCTGTCTTCATCTTC | O ATGTACTCATGATCTCCCTGAGCCCCATAGTCACATGTGTGGTGGTGGTGGTGTGTGGGGGGGG | 5 AIGIGGLICACCATIACICIGACICCIAAGGICACGIGIGIGIGGIG | A THE NEW TO CHARACTER THE THE THE THE TABLE TO CHARACTER THE THE THE THE THE THE THE THE THE THE | 0 CACAAACCCATAGAGGATTACAACAGTACTCTCCGGGTGGTC | GCAACCCCGGGAGGAGCAGTTCAACAGCACTTTCCGCTCAGTCAG | AGCACCAGGACTGGATGACAAGAGTTCAAATGCAAGGTCAACAACAACAAAGACCTCC |    |      | O AIGTCTIGCCTCCACCAGAAGAAGAGATGACTAAGAAACAGGTCACTCTGACCTGCATGG | 3 ACACCATTCCACCACCAGGGGGGGGGGGGGGGGGGGGGG | 0 TCACAGACTICATGCCTGAAGACATTTACGTGGAGTGGACCAACAGGGAAAACAGGC | 3 taacadacritcriccridaadacattacristigadridecagriga | 0 TATACTACAAGAACACTGAACCCTGGACTCTGATGGTTCTTACTTCATGTACAGCA | 3 AGAACTACAAGAACACTCAGCCCATCATGGACACAGATGGCTCT | O AGCTGAGAGAGAAAGAAGAACTGGGTGGAAAGAAATAGCTACTCCTGTTCAGTGGTCC |      |          | 6 standard; DNA; 1045 BP.      | 9       | 2004 (first entry) | protein WT-IL-15-wlgG2a coding sequence. | immunosuppressive; antirheumatic; antiarthritic; antid:<br>neuroprotective; antipsoriatic; dermatological; antiin:<br>cytostatic; interleukin-15; immunoglobulin G; ds; gene | ic.<br>1fled.        | Location/Qualifiers 11044 /*teg= a /product= "WT-IL-15-mlgG2a" /transl except |
| 155:                                            | 79.                                           | 161                  | 83                                                      | 167                                            | σ .                                                                | 1/3                                               | ח ה                                                                                               | 101                                          | 185.                                          | 1070                                                       |    | 1973 | 119                                                            | 203                                       | 125(                                                        | 209                                                | 131(                                                       | 215                                            | 137(                                                         | 43   | 27       | 3756                           | AD00756 | 15-JUL-3           | Fusion p                                 | immunosu<br>neuropro<br>cytostat                                                                                                                                             | Syntheti<br>Unidenti | Key<br>CDS                                                                    |
| g                                               | 8                                             | Q                    | à                                                       | qq                                             | රි සි                                                              | 8 8                                               | <b>3</b> 8                                                                                        | È                                            | අු                                            | රි සි                                                      | ઠે | 음    | ò                                                              | QQ                                        | ò                                                           | qq                                                 | È                                                          | <b>q</b>                                       | <b>∂</b> 8                                                   | 8    | <b>a</b> | RESULT 8<br>ADO07566<br>ID ADO | ¥ ¥     | XE:                | XB;                                      | \$ & & & ;                                                                                                                                                                   | វុខខ                 | 3                                                                             |

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976 rcaeregrecaceaceacererecacacaceaceaceaceaceaceacereceeeerece 1035
   New fusion protein of interleukin-15 and Pc fragment, useful for treating e.g. transplantation disorders, autoimmune diseases and tumors, also related nucleic acid.
   The present invention relates to a fusion protein consisting of wild-type interleukin-15 (IL-15) and an immunoglobulin G (IgG) Fc fragment, other than a murine IgG2b Fc fragment. The fusion proteins and coding sequences are used to prevent or treat consequences of transplantation and/or autoimmune diseases, e.g. rhematoid arthritis, diabetes, multiple sclerosis, psoriasis, neurodermatitis, ulcerative colitis, tumours and AIDS, etc., and tissues or organs that express the protein are useful for
   1420 TCAGTGGTCCACGAGGTCTCCACAATCACCACGACTAAGAGCTTCTCCCGGACTCCG
                        1180 CCACAGGIATATGICTTGCCTCCACCAGAAGAGAGAGATGACTAAGAAACAGGTCACTCTG
  1240 ACCTGCATGGTCACAGACTTCATGCCTGAAGACATTTACGTGGAGTGGACCAACAAGGG
  1300 AAAACAGAGCTAAAACTACAAGAACACTGAACCAGTCCTGGACTCTGATGGTTCTTACTTC
  736 ccacaggrarargrerrecerceacagaagaagagargacraagaaacaggreaerere
   immunosuppressive; antirheumatic; antiarthritic; antidiabetic; neuroprotective; antipsoriatic; dermatological; antiinflammatory; cytostatic; interleukin-15; immunoglobulin G; ds; gene; human.
  Fusion protein coding sequence fragment 149-Fc.
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  Disclosure; Fig 11; 63pp; German.
  BP.
   (HOFF ) HOFFMANN LA ROCHE & CO
  AD007578 standard; DNA; 1108
  13-OCT-2003; 2003WO-CH000666
   14-OCT-2002; 2002EP-00022869
  (first entry)
  1480 GGTAAA 1485
  GGTAAA 1041
  WPI; 2004-357203/33.
  Dreher I, Moll
  WO2004035622-A2.
  Synthetic.
Unidentified.
  29-APR-2004.
  15-JUL-2004
      1120
  AD007578;
  1036
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  1059
  CTCCCCATCCAGCACCAGGACTGGATGAGTGGCAAGGAGTTCAAATGCAAGGTCAACAAC 1119
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  The present invention relates to a fusion protein consisting of wild-type interleukin-15 (IL-15) and an immunoglobulin G (IgG) Fc fragment, other than a murine IgG2b Fc fragment. The fusion proteins and coding sequences are used to prevent or treat consequences of transplantation and/or autoimmune diseases, e.g. rheumatoid arthritis, diabetes, multiple sclerosis, psoriasis, neurodermatitis, ulocative colitis, tumours and AIDS, etc., and tissues or organs that express the protein are useful for transplantation into humans or other mammals, as allo, auto- or xenotransplants. Also transgenic animals that express the fusion proteins are useful as source of cells, tissues and organs for transplantation or to screen for pharmaceuticals and/or to identify toxic substances. The present sequence is a polypeptide coding sequence used in the
  819
  759
  675
  375
  879
  435
  939
  999
  555
  615
  treating
  700 AAAGAGCTCCAGGCCCTGGAGAAAGGAAAATGCACAGCTGGAATGGGAGTTGCAAGCACTG
   316 ATTGTCCAAATGTTCATCAACACTTCGGATCCCAGAGGGCCCACAATCAAGCCCTGTCCT
   376 CCATGCAAATGCCCAGCACCTAACCTCTTGGGTGGACCATCCGTCTTCATCTTCCCTCCA
  616 CTCCCCATCCAGCACCAGGACTGGATGAGGAGTTCAAATGCAAGGTCAAACAAC
  256 AAAGAATGTGAGGAACTGGAGGAAAAAAATATTAAAGAATTTTTGCAGGGTTTTTGTACAT
  760 GAAAAGGAACTGGCTCAGGCAGCATCTGAGCCCCAGAGGCCCCACAATCAAGCCCTGTCCT
  CCATGCAAATGCCCAGCACCTAACCTCTTGGGTGGACCATCCGTCTTCATCTTCCCTCCA
  880 AAGAICAAGGAIGIACTCATGAICTCCCTGAGCCCCATAGTCACAIGIGGGGGGGAI
  436 AAGATCAAGGATGTACTCATGATCTCCCTGAGCCCCCATAGTCACATGTGTGGTGGTGGTGT
  GTGAGCGAGGATGACCCAGATGTCCAGATCAGCTGGTTTGTGAACAACGTGGAAGTACAC
  496 Greadceadeareacceaearerceaeareaceaerrearreaeacaaceaecreeaecae
  ACAGCTCAGACACAAACCCATAGAGAGGATTACAACAGTACTCCCGGGTGGTCAGTGCC
  556 ACAGCTCAGACACAAACCCATAGAGGATTACAACAGTACTCTCCGGGTGGTCAGTGCC
  new fusion protein of interleukin-15 and Pc fragment, useful for tres
e.g. transplantation disorders, autoimmune diseases and tumors, also
related nucleic acid.
  Gaps
   / Match 47.2%; Score 701.2; DB 12; Length 1045; Local Similarity 93.3%; Pred. No. 1.6e-178; nes 733; Conservative 0; Mismatches 53; Indels 0:
  Seguence 1045 BP; 331 A; 238 C; 235 G; 241 T; 0 U; 0 Other;
        Thr"
             aa:
/note= "(pos: 34. .36, /partial
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  exemplification of the invention.
  /note= "no start
  Disclosure; Fig 5; 63pp; German.
  (HOFF ) HOFFMANN LA ROCHE & CO
  13-OCT-2003; 2003WO-CH000666.
  14-OCT-2002; 2002EP-00022869.
  WPI; 2004-357203/33.
  Dreher I, Moll T;
  P-PSDB; AD007559
  WO2004035622-A2
  29-APR-2004
  1000
  820
  940
  ery Match
  st Loca
tches
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transplantation into humans or other mammals, as allo-, auto- or xeno-
transplants. Also transpenic animals that express the fusion proteins are
useful as source of cells, tissues and organs for transplantation or to
screen for pharmaceuticals and/or to identify toxic substances. The
present sequence is a coding sequence used in the exemplification of the
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AD007577;

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Sequence 1108 BP; 343 A; 255 C; 253 G; 257 T; 0 U; 0 Other;
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  1059
  1119
  1179
   1239
   1299
   1359
   1419
   1038
   1479
  1098
   378
   819
   879
  678
  738
  798
   828
   918
   978
   700 AAAGAGCTCCAGGCCCTGGAGAAAAATGCACAGCTGGAATGGGAGTTGCAAGCACTG 759
   438
   498
   939
  558
   CCATGCAAATGCCCAGCACCTCTTGGGTGGACCATCCGTCTTCATCTTCCCTCCA
   CCATGCAAATGCCCAGCACCTAACCTCTTGGGTGGACCATCCGTCTTCATCTTCCCTCCA
   AAGATCAAGGATGTACTCATGATCTCCCTGAGCCCCATAGTCACATGTGTGGTGGTGGTTGTT
   Greadecadearcacadarcrecadarcacerecrecimienteaacaacerecaagracae
  ACAGCTCAGACACAAACCCATAGAGAGGATTACAACAGTACTCTCCGGGTGGTCAGTGCC
  CTCCCCATCCAGCACCAGGACTGGATGAGTGGCAAGGAGTTCAAATGCAAGGTCAACAAC
  CTCCCCATCCAGCACCAGGATGAGTGGCAAGGAGTTCAAATGCAAGGTCAACAAC
  AAAGACCTCCCAGCGCCCATCGAGAGAACCATCTCAAAACCCAAAGGGTCAGTAAGAGCT
   CCACAGGTATATGTCTTGCCTCCACCAGAAGAAGAGATGACTAAGAAACAGGTCACTCTG
   CCACAGGTATATGTCTTGCCTCCACCAGAAGAAGAGAGACTAAGAAACAGGTCACTCTG
   ACCTGCATGGTCACAGACTTCATGCCTGAAGACATTTACGTGGAGTGGACCAACAACGGG
   1039 TCAGTGGTCCACGAGGGTCTGCACAATCACCACACGACTAAAGAGCTTCTCCCGGAGTCCCG
   GAAAAGGAACTGGCTCAGGCAGCATCTGAGCCCCAGAGGGCCCACAATCAAGCCCTGTCCT
   379 ATTGTCCAAATGTTCATCAACACTTCGGATCCCAGAGGGCCCACAATCAAGCCCTGTCCT
  GTGAGCGAGGATGACCCAGATGTCCAGATCAGCTGGTTTTGTGAACAACGTGGAAGTACAC
  ACAGCTCAGACACAAACCCATAGAGAGGATTACAACAGTACTCTCCGGGTGGTCAGTGCC
  AAAGACCTCCCAGGGCCCCATCGAGAGAACCATCTCAAAAACCCAAAGGGTCAGTAAGAGCT
   1300 AAAACAGAGCTAAAACTACAAGAACACTGAAACCAGTCCTGGACTCTGATGGTTCTTACTTC
   AAAACAGAGCTAAAACTACAAGAACACTGAACCAGTCCTGGACTCTGATGGTTCTTACTTC
   TCAGTGGTCCACGAGGGTCTGCACAATCACCACACGACTAAGAGCTTCTCCCGGACTCCG
   Gaps
uery Match 47.2%; Score 701.2; DB 12; Length 1108; set Local Similarity 93.3%; Pred. No. 1.7e-178; atches 733; Conservative 0; Mismatches 53; Indels 0;
  GGTAAA 1485
  GGTAAA 1104
   260
  1000
  619
  1060
  619
  1120
   1180
   799
   1240
   859
   919
   1360
   1420
  1480
  1099
   820
   439
   880
  499
  940
  739
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The present invention relates to a fusion protein consisting of wild-type interleukin-15 (IL-15) and an immunoglobulin G (IgG) Pc fragment, other than a murine IgG2D Pc fragment. The fusion proteins and coding sequences are used to prevent or treat consequences of transplantation and/or autoimmune diseases, e.g. rheumatoid arthritis, diabetes, multiple sclerosis, psoriasis, neurodermatitis, ulcerative colitis, tumours and AIDS, etc., and tissues or organs that express the protein are useful for transplantation into humans or other mammals, as allo-, auto- or xeno-transplants. Also transpenic animals that express the fusion proteins are useful as source of cells, tissues and organs for transplantation or to screen for pharmaceuticals and/or to identify toxic substances. The present sequence is a coding sequence used in the exemplification of the
  New fusion protein of interleukin-15 and Pc fragment, useful for treating e.g. transplantation disorders, autoimmune diseases and tumors, also related nucleic acid.
  319 AAAGAATGTGAGGAACTGGAGGAAAAAAAATATTAAAGAATTTTTGGACAGTTTTGTACAT 378
  CCATGCAAATGCCCCAGCACCTAACCTCTTGGGTGGAACCATCCGTCTTCATCTTCCCTCCA
   AAGATCAAGGATGTACTCATGATCTCCCTGAGCCCCATAGTCACATGTGTGGTGGTGGTGGAT
  GTGAGGAGGATGACCCAGATGTCCAGATCAGCTGGTTTGTGAACAACGTGGAAGTACAC
  700 AAAGAGCTCCAGGCCCTGGAGAAGGAAATGCACAGCTGGAATGGGAGTTGCAAGCACTG
   760 GAAAAGGAACTGGCTCAGGCATCTGAGCCCAGAGGGCCCACAATCAAGCCCTGTCCT
   CCATGCAAATGCCCAGCACCTAAACCTCTTGGGTGGACCATCCGTCTTCATCTTCCCTCCA
  AAGATCAAGGATGTACTCATGATCTCCCTGAGCCCCATAGTCACATGTGGTGGTGGTGGTGGT
   GTGAGCGAGCATGACCCAGATGTCCAGATCAGCTGGTTTGTGAACAACGTGGAAGTACAC
  immunosuppressive, antirheumatic, antiarthritic; antidiabetic; neuroprotective; antipsoriatic; dermatological; antiinflammatory; cytostatic; interleukin-15; immunoglobulin G; ds; gene.
  Length 1108;
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   Sequence 1108 BP; 342 A; 255 C; 254 G; 257 T; 0 U; 0 Other;
   Score 701.2; DB 12; Length.
Pred. No. 1.7e-178;
0; Mismatches 53; Indels
  Pusion protein coding sequence fragment
   ğ
   Disclosure, Fig 10; 63pp; German.
   8
   14-OCT-2002; 2002EP-00022869.
   unery Match
Best Local Similarity 93.3%;
Matches 733; Conservative
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  (first entry)
  WPI; 2004-357203/33.
   (HOFF ) HOFFMANN LA
  Moll T;
  WO2004035622-A2
   13-OCT-2003;
   Synthetic.
Unidentified
  15-JUL-2004
   29-APR-2004
  Dreher I,
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AD007577 standard; DNA; 1108

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   1098
  1179
   1239
   738
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  828
   918
   978
  mouse; murine; antibody; gene; ds; beta-amyloid; A-beta; amyloid beta A4 precursor protein; APP; presenilin; lipoprotein receptor related protein; LRP; beta-amyloid 42; A-beta 42; Alzheimer's disease; neuroprotective; nootropic.
CCACAGGTATATGTCTTGCCTCCACAGAAGAAGAGATGACTAAGAAACAGGTCACTCTG
  1420 TCAGTGGTCCACGAGGGTCTGCACAATCACCACACGACTAAAGAGCTTCTCCCGGACTCCG
  AAAGACCTCCCAGCGCCCATCGAGAGAACCATCTCAAAACCCAAAGGGTCAGTAAGAGCT
  739 AAAGACCTCCCAGCGCCCATCGAGAGCCATCTCAAAACCCAAAGGGTCAGTAAGAGCT
   CCACAGGTATATGTCTTGCCTCCACCAGAAGAGAGATGACTAAGAAACAGGTCACTCTG
  1240 ACCTGCATGGTCACAGACTTCATGCCTGAAGACATTTACGTGGAGTGGACCAACAACGG
   859 ACCTGCATGGTCACAGACTTCATGCCTGAAGACATTTACGTGGAGTGGACCAACAACGGG
  AAAACAGAGCTAAAACTACAAGAACACTGAACCAGTCCTGGACTCTGATGGTTCTTACTTC
   Murine immunoglobulin heavy chain constant region DNA SeqID
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   Wagner
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   Velicelebi
   BP
   20-AUG-2002; 2002US-0405417P.
18-SEP-2002; 2002US-0411974P.
  20-AUG-2003; 2003WO-US026173
   ADL15694 standard; DNA; 990
  (first entry)
   Kounnas M, Patrick A,
  (NEUR-) NEUROGENETICS
  GGTAAA 1485
  GGTAAA 1104
  WPI; 2004-226902/21.
   P-PSDB; ADL15695
  WO2004018997-A2
   Mus musculus.
  20-MAY-2004
   04-MAR-2004
   1000
  799
   1300
   979
  1099
  1060
   1120
   1180
   1480
   ADL15694;
```

New polypeptide comprises a sequence of amino acids that is selectively reactive with beta-amyloid peptide 42 or at least one complementarity-

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This invention relates to novel methods and compositions for detecting and modulating beta-amyloid (A-beta) peptide levels and the processing of amyloid beta A precursor protein (APP). Specifically, it refers to methods of assessing the presentlin activity of compounds using the lipoprotein receptor related protein (LRP), in order to identify to presentlin proteins that can be used to affect the processing of APP. The presentlin proteins that can be used to affect the processing of APP. The presentlin activity and A-beta levels, in particular beta-amyloid 42 (A-beta 42), such that the agent is selectivly reactive with A-beta 42 and binds at least one complementarity determining region (CDR) of either antibody A387 or antibody 8436. As such, the polypeptides, nucleic acids and antibodies are useful for treating Alzheimer's disease, accordingly the compositions exhibit neuroprotective and nootropic activities. This invention.
  1085
  1145
  1325
   1445
  GATCAGCTGGTTTGTGAACAACGTGGAAGTACACACAGCTCAGACACAAACCCATAGAGA 1025
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   410
  590
   950
  845
   470
   905
   965
  GAAGAACTGGGTGGAAAGAAATAGCTACTCCTGTTCAGTGGTCCACGAGGGTCTGCACAAA
  GAGTGGCAAGGAGTTCAAATGCAAGGTCAACAACAACACCTCCCAGCGCCCATCGAGAG
  AGAAGAAGAAGATGACTAAGAAACAGGTCACTCTGACCTGCATGGTCACAGACTTCATGCC
   TGAAGACATTTACGTGGAGTGGACCAACAACGGGAAAACAGAGCTAAAACTACAAGAACAC
   GGATTACAACAGTACTCTCCGGGTGGTCGTGCCCTCCCCCATCCAGCACCAGGA
   1206 AGAAGAAGAGATGACTAAGAAACAGGTCACTGACCTGCATGGTCACAGACTTCATGCC
  TGAACCAGTCCTGGACTCTGATGGTTCTTACTTCATGTACAGCAGAGGTGGGAAAA
  CTTGGGTGGACCATCCGTCTTCATCTTCCCTCCAAAGATCAAGGATGTACTCATGATCTC
  CTTGGGTGGACCATCCGTCTTCATCTTCCTCCAAAGATCAAGGATGTACTCATGATCTC
  GGATTACAACAGTACTCTCCGGGTGGTCAGTGCCCTCCCCATCCAGCACCAGGACTGGAT
   AACCATCTCAAAACCCAAAGGGTCAGTAAGAGCTCCACAGGTATATGTCTTGCCTCCACC
  TGAGCCCAGAGGCCCACACATCAAGCCCTGTCCTCCATGCAAATGCCCAGCACCTCTAACCT
   291 TGAGCCCAGAGGGCCCACAATCAAGCCCTGTCCTCCATGCAAATGCCCAGGCACCTAACCT
  CCTGAGCCCCATAGTCACATGTGTGGTGGTGTGTGAGCGAGGATGACCCAGATGTCCA
   Gaps
   ö
   47.2%; Score 700.6; DB 12; Length 990; 100.0%; Pred. No. 2.3e-178; arive 0; Mismatches 0; Indels 0;
   Sequence 990 BP; 274 A; 286 C; 235 G; 195 T; 0 U; 0 Other;
  TCACCACACGACTAAGAGCTTCTCCCGGAGTCCCGGGTAAA 1485
region of antibody A387 or B436, useful disease.
   Disclosure; SEQ ID NO 68; 408pp; English
   Best Local Similarity 100.
Matches 700; Conservative
       determining
Alzheimer's
   786
  351
   1026
  531
   1086
   1146
  651
  711
   1266
   771
   831
  1386
  891
  1446
   996
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951 TCACCACACGACTAAGAGCTTCTCCCGGACTCCGGGTAAA 990
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antitheumatic; antithyroid; bon metastases; calcium antagonist; cancer; cardiovascular-gen.; degeneration; eating-disorders-gen.; cardiovascular-gen.; endocrine-gen.; gastrointestinal-gen.; gene; genetic disorder; heavy chain; hepatotropic; hypercalcenta; immune disorder; heavy chain; hepatotropic; hypercalcenta; outh disease; musculoskeletal disease; moplaem, nephrotropic; osteopethic; osteopethics; osteopethics; pagets disease; periodontal disease; pharmaceutical; rheumatold arthritis; ss. endocrine-gen.; antiarthritic; antibacterial; antiinflammatory; M-CSF specific murine antibody RX1 heavy chain cDNA. AEC20762 standard; cDNA; 1401 20-OCT-2005 (first entry) AEC20762;

Mus musculus.

WO2005068503-A2.

28-JUL-2005

06-JAN-2005; 2005WO-US000546

07-JAN-2004; 2004US-0535181P. 02-JUN-2004; 2004US-0576417P.

(CHIR ) CHIRON CORP. (XOMA ) XOMA TECHNOLOGY LTD.

Liu C, Zimmerman DL, Harrowe GM, Koths K, Kavanaugh WM, Long L; Calderon-Cacia M, Horwitz AH;

2005-597707/61. P-PSDB; AEC20763

for Novel non-murine antibody that competes with monoclonal antibody RXI binding to macrophage colony stimulating factor, useful for treating hypogonadism, hypercalcemia, rickets, scurvy, homocystinuria, cancer, osteoporosis.

Claim 67; SEQ ID NO 1; 269pp; English.

The invention describes a non-murine antibody (I) that competes with monoclonal antibody RX1 for binding to macrophage colony stimulating factor (M-CSF) by more than 75¢, where the monoclonal antibody RX1 has the heavy chain and light chain amino acid sequences having a fully defined 447 amino acids (SEQ ID No. 2) and 214 amino acids (SEQ ID No. 4) sequences given in the specification, respectively. (I) is useful for preventing a subject afflicted with a disease that causes or contributes to osteolysis, where the antibody effectively reduces the severity of bone loss associated with the disease. The disease is chosen from metabolic bone diseases associated with relatively increased osteoclast activity, including endocrinopathies, hypercalcemia, deficiency states, chronic diseases, and hereditary diseases, cancer, osteoporosis, osteopetrosis, inflammation of bone associated with arthritis and rheumatoid arthritis, periodontal disease, fibrous dysplasia, and/or paget's disease. (I) is useful for preventing or treating metastatic cancer. Antibodies of the invention are useful for preventing or reducing bone loss; osteolysis; metastatic cancer to bone and cancer. (I) is useful for seventing or reducing bone loss in a patient exhibiting osteolysis, manufacturing a medicament for preventing a patient for mentastatic cancer to bone in a patient suffering from metastatic cancer. (I) in synergistic combination, is useful for preparing a patient having cancer. (I) in synergistic combination, is useful for preparing a

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1205
  1121
  1025
   1085
   1086 GAGTGGCAAGGAGTTCAAATGCAAGGTCAACAACAAAGACCTCCCAGCGCCCATCGAGAG 1145
   1206 AGAAGAAAGAGATGACTAAGAAACAGGTCACTGTGACCTGCATGGTCACAGACTTCATGCC 1265
   1122 AGAAGAAGAAGATGACTAAGAAACAGGTCACTCTGACCTGCATGGTCACAGACTTCATGCC 1181
   1182 TGAAGACATTTACGTGGAGTGGACCAACAACGGGAAAACAGGGCTAAACTACAAGAACAC 1241
   1386 GAAGAACTGGGTGGAAAGAAATAGCTACTCCTGTTCAGTGGTCCACGAGGGTCTGCACAA 1445
  1001
  ö
  845
  761
  965
  941
   821
  822 cereadececcaragreacarereresesesesarereadeceadareaceadareree 881
   cerebroprotective; cell therapy; gene therapy; drug delivery; prion disease; 44B1H; ds; therapeutic; monoclonal antibody; heavy chain.
medicament for treating a patient exhibiting osteolysis. This sequence encodes macrophage colony stimulating factor (M-CSF) specific murine antibody RX1 heavy chain.
  942 GGATTACAACAGTACTCTCCGGGTGGTCAGTGCCCTCCCCCATCCAGCACCAGGACTGGAT
   1062 AACCATCTCAAAAACCCAAAGGGTCAGTAAGAGCTCCACAGGTATATGTCTTGCCTCCACC
   762 criederedecerreceretrearcricereceaagareaagargraetergarete
   1026 GGATTACAACAGTACTCTCCGGGTGGTCAGTGCCCTCCCCATCCAGGACCAGGACTGGAT
  1146 AACCATCTCAAAACCCAAAGGGTCAGTAAGAGCTCCACACAGGTATATGTCTTGCCTCCACC
   TGAAGACATTTACGTGGAGTGGACCAACAACGGGAAAACAGAGCTAAAACTACAAGAACAC
  1326 TGAACCAGTCCTGGACTCTGATGGTTCTTACTTCATGTACAGCAAGCTGAGAGAAAA
  786 TGAGCCCAGAGGCCCAGATCAAGCCCTGTCCTCCATGCAAATGCCCCAGCACCTAACCT
   702 TGAGCCCAGAGGCCCACAATCAAGCCCTGTCCTCCATGCAAATGCCCAGCACCTAACCT
   CTTGGGTGGACCATCCGTCTTCATCTTCCCTCCAAAGATCAAGGATGTACTCATGATCTC
  906 CCTGAGCCCCATAGTCACATGTGTGTGGTGGATGTGAGCGAGGATGACCCAGATGTCCA
  GATCAGCTGGTTTGTGAACAACGTGGAAGTACACACACAGCTCAGACACAAACCCATAGAGA
  Gapa
  Length 1401;
  ö
   Sequence 1401 BP; 369 A; 407 C; 327 G; 298 T; 0 U; 0 Other;
  0; Indels
  1362 TCACCACCACGACTAAGAGCTTCTCCCGGACTCCGGGTAAA 1401
   1446 TCACCACACGACTAAGAGCTTCTCCCGGACTCCGGGTAAA 1485
  Anti-PrP antibody heavy chain 44BlH DNA SEQ ID NO 5.
   Query Match
47.2%; Score 700.6; DB 14;
Best Local Similarity 100.0%; Pred. No. 2.7e-178;
Matches 700; Conservative 0; Mismatches 0;
  AED19725 standard; DNA; 1560 BP
  15-DEC-2005 (first entry)
  Unidentified
   846
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  996
  AED19725;
   RESULT 13
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1090 GGATTACAACAGTACTCTCCGGGTGGTCAGTGCCCTCCCCATCCAGCACCAGGACTGGAT 1149
   1450 GAAGAACTGGGTGGAAAGAAATAGCTACTCCTGTTCAGTGGTCCACGAGGGTCTGCACAA 1509
CCTGAGCCCCATAGTCACATGTGTGGTGGTGGATGTGAGCGAGGATGACCCAGATGTCCA 1029
  1330 TGAAGACATTTACGTGGAGTGGACCAACACGGGAAAACAGGGAAAACTAAACTACAAGAACAC 1389
   1086 GAGTGGCAAGGAGTTCAAATGCAAGGTCAACAACAAGACCTCCCAGCGCCCCATCGAGAG
  1146 AACCATCTCAAAACCCAAAGGGTCAGTAAGAGCTCCACAGGTATATGTCTTGCCTCCACC
   1390 reaaccagrecresacrerearesrecreaterrerrearescasseas
   GATCAGCTGGTTTGTGAACAACGTGGAAGTACACACAGGCTCAGACACAACCCATAGAGA
   GGATTACAACAGTACTCTCCGGGTGGTCAGTGCCCTCCCCATCCAGCACCAGGACTGGAT
   1206 AGAAGAAAGAGTAAGAAACAGGTCACTGTGACCTGCATGGTCACAGACTTCATGCC
  1266 TGAAGACATTTACGTGGAGTGGACCAACAACGGGAAAACAGAGCTAAACTACAAGAACAC
   TGAACCAGTCCTGGACTCTTGTTCTTTCATGTACAGCAAGCTGAGAAAA
   1386 GAAGAACTGGGTGGAAAGAAATAGCTACTCCTGTTCAGTGGTCCACGAGGGTCTGCACAA
  New de-immunized anti-CD3 antibody, useful for stimulating an immune response against infections and for treating infections.
   ds; gene; immunostimulant; immunogenicity; antibody.
   Carr FJ,
   'n
  Disclosure, Fig 1a, 75pp; English.
   3
  ADV26108 standard; DNA; 1569
   28-MAY-2004; 2004WO-US017219.
   02-JUN-2003; 2003US-0475155P.
   Rother RP, Faas-Knight S,
  10-MAR-2005 (first entry)
  (ALEX-) ALEXION PHARM INC.
  WPI; 2005-031597/03.
  Mouse OKT3 VH gene.
  P-PSDB; ADV26107.
   WO2004108158-A1.
  16-DEC-2004.
  996
  1026
  1326
   ADV26108;
                                970
  Mus sp.
  RESULT 14
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   The invention describes an agent (I) for treating prion disease or delivering a subtance to the lesioned region of prion disease, comprising a mesenchymal cell. Also described are: a nucleic acid (II) having an anti-prion antibody gene comprising: an antibody heavy chain gene having SEO ID No: 1, 3, 5, 30, 32 and 34, an unclectide sequence consisting of a degenerate genetic code, which encodes a polypeptide same as that of the above sequences, or a nucleotide sequence, which is complementary to the above sequences; and an antibody light chain gene having SEO ID No: 2, 4, 6, 31, 33 and 35, a nucleotide sequence furtis consisting of degenerate genetic code, which encodes a polypeptide same as that of the above nucleotide sequence and that hybridizes under stringent consisting of degenerate genetic code, which encodes a polypeptide same as that of the above nucleotide sequence and that hybridizes under stringent complementary to the above sequences, or a nucleotide sequence that is complementary to the above sequences, a nucleotide sequence that is encoded by (II) and consistant region of antibody of antibody of antibody encoded by (II) and constant region of antibody of antibod other than mouse; a nucleic acid that encodes (IV); preparing (II) and constant region of antibody of antibody of antibody of antibody of antibody of antibody of antibody of antibody of antibody of antibody of antibody utilized for treating prion proliferation inhibition activity, to acid; that encodes (IV); preparing (II) or (III); a cell (V) having abnormal prion proliferation inhibition activity, occupation of a prion disease; and approximate antibody mescander control of a prion disease; and delivering a substance to the lesioned region of prion disease; and delivering a substance to the lesioned region of prion disease; and delivering a substance to the lesioned region of prion disease; and delivering a substance to the lesioned region of prion disease; and delivering a substance to the lesioned region of prion disease; and delivering
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  845
  905
  696
   850 TGAGCCCAGAGGCCCACAATCAAGCCCTGTCCTCCATGCAAATGCCCAGCACCTAACCT 909
   906 CCTGAGCCCCATAGTCACATGTGGTGGTGGATGTGAGCGAGGATGACCCAGATGTCCA 965
  Agent useful for treating prion disease or delivering a substance to a
lesioned region of prion disease, comprises a mesenchymal cell.
  846 CTTGGGTGGACCATCCTTCATCTTCCCTCCAAAGATCAAGGATGTACTCATGATCTC
  910 CTTGGGTGGACCATCCGTCTTCATCTTCCCTCCAAAGATCAAGGATGTACTCATGATGTCTC
  TGAGCCCAGAGGGCCCACAATCAAGCCCTGTCCTCCATGCAAATGCCCAGCACCTAACCT
  0; Сарв
   ery Match 47.2%; Score 700.6; DB 14; Length 1560; Bt Local Similarity 100.0%; Pred. No. 2.8e-178; tches 700; Conservative 0; Mismatches 0; Indels 0;
  Ë
   Sequence 1560 BP; 413 A; 424 C; 388 G; 335 T; 0 U; 0 Other;
  Hamada H, Horiuchi
   Fujinaga K, Shinagawa M, Niitsu Y,
Honmou O, Umetani A;
   Claim 4; SEQ ID NO 5; 34pp; Japanese.
   30-MAR-2005; 2005WO-JP006189.
  30-MAR-2004; 2004JP-00100649
   (RENO-) RENOMEDIX INST INC
  WPI; 2005-725409/74.
                                  WO2005094846-A1.
   13-0CT-2005.
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Hamilton A;

28-MAY-2004; 2004WO-US017005

WO2005007809-A2 27-JAN-2005.

Mus sp. Synthetic.

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1265
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  966 GATCAGCTGGTTTGTGAACAACGTGGAAGTACACACAGCTCAGACACAAACCCATAGAGA 1025
   1085
   1043
  1145
   1103
  1205
  1163
   1164 AGAAGAAGAGATGACTAAGAAACAGGTCACTCTGACCTGCATGGTCACAGACTTCATGCC 1223
  1385
  1386 GAAGAACTGGGTGGAAAGAAATAGCTACTCCTGTTCAGTGGTCCCACGAGGGTCTGCACAA 1445
   1344 GAAGAACTGGGTGGAAAGAAATAGCTACTCCTGTTCAGTGGTCCACGAGGGTCTGCACAA 1403
  845
   803
  905
   863
  965
   923
   983
anti-CD3 antibody. The antibody,
  744 TGAGCCCAGAGGGCCCACAATCAAGCCCTGTCCTCCATGCAAATGCCCAGCACCTAACCT
   804 CTTGGGTGGACCATCCGTCTTCATCTTCCCTCCAAAGATCAAGGATGTACTCATGATCTC
   906 CCTGAGCCCCATAGTCACATGTGTGGGTGGATGTGAGCGAGGATGACCCAGATGTCCA
  864 CCTGAGCCCCATAGTCACATGTGTGGTGGATGTGAGCGAGGATGATGACCCAGATGTCCA
  924 GATCAGCTGGTTTGTGAACAACGTGGAAGTACACACAGGTCAGACACAAACCCATAGAGA
   1026 GGATTACAACAGTACTCTCCGGGTGGTCAGTGCCCTCCCCATCCAGCACCAGGACTGGAT
   1086 GAGTGGCAAGGAGTTCAAATGCAAGGTCAACAAAAAAACACCTCCCAGCGCCCATCGAGAG
   104 AACCATCTCAAAACCCAAAAGGGTCAGTAAGAGCTCCACAGGTATATGTCTTGCCTCCACC
   AGAAGAAGAGAGAGACTAAGAAACAGGTCACTCTGACCTGCATGGTCACAGACTTCATGCC
   TGAAGACATTTACGTGGAGTGGACCAACAACGGGAAAACAGAGCTAAAGTACAAGAACAC
   TGAACCAGTCCTGGACTCTGATGGTTCTTACTTCATGTACAGCAAGCTGAGAGTGGAAAA
  846 CTTGGGTGGACCATCCGTCTTCATCTTCCCTCCAAAGATCAAGGATGTACTCATGATCTC
  984 GGATTACAACAGTACTCTCCGGGTGGTCAGTGCCCTCCCCATCCAGCACCAGGACTGGAT
  1146 AACCATCTCAAAACCCAAAAGGGTCAGTAAGAGCTCCACAGGTATATGTCTTGCCTCCACC
The invention relates to a de-immunized anti-CD3 antibody. The antiboc composition and methods are useful for stimulating an immune response against infections and for treating infections. The present sequence represents DNA encoding mouse OKT3 VH.
   786 TGAGCCCAGAGGCCCCACAATCAAGCCCTGTCCTCCATGCAAATGCCCAGCACCTAACCT
   Gaps
   uery Match 47.2%; Score 700.6; DB 14; Length 1569; isst Local Similarity 100.0%; Pred. No. 2.8e-178; atches 700; Conservative 0; Mismatches 0; Indels 0;
   Sequence 1569 BP; 446 A; 443 C; 356 G; 324 T; 0 U; 0 Other;
  TCACCACACACTAAGAGCTTCTCCCGGACTCCGGGTAAA 1485
   1206
   1326
   1446
   1266
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   fusion protein; IgG; antibody; antiinflammatory; antimicrobial-gen.; cytokine release syndrome; inflammation; infection; ds; gene.
   heavy chain variable region DNA.
                               ВР
                               ADW71834 standard; DNA; 1569
  (first entry)
  07-APR-2005
  Murine OKT3
  ADW71834;
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The invention relates to a novel fusion protein, comprising a non-FC component and an FC region having a first portion derived from one or more human igg2 antibodies and a second portion derived from one or more human igg2 antibodies. The invention further comprises: a method for reducing antibodies. The invention further comprises: a method for comprising administering an antibody which binds to either a cell surface molecule or a soluble molecule that binds to a cell surface molecule or a soluble molecule that binds to acli surface molecule or a soluble molecule that binds to acli surface molecule or a more human igg2 antibodies and a second continuod derived from one or more human igg2 antibodies and a second continuod derived from one or more human igg2 antibodies and a second continuod derived from one or more human igg2 antibodies and a second continuod from one or more human igg2 antibodies and a second component for a mon-Fc component increasing the avidity of component for a mon-Fc component in mammalian cells creceptor binding adder of a non-Fc component in mammalian cells in protein, or improving expression of a non-Fc component in mammalian cells in a nucleic acid encoding a fusion protein, an expression vector containing the nucleic acid; a host cell transfected with an expression vector; and a composition comprising the fusion protein and a preventing or reducing antibody-mediated cell activation or preventing or reducing cytokine release or the severity of cytokine release syndrome, and in reducing antibody-mediated cell activation or inflammation events. The antibody-mediated cell activation or composition, methods are useful for preventing of the invention. This
   GATCAGCTGGTTTGTGAACAACGTGGAAGTACACAGGCTCAGACACAAACCCATAGAGA 1025
   845
  863
   864 ccricadececearadereardrefredesereardregesereardaeceadardrefea 923
   Reducing antibody-mediated cell activation or inflammation events, useful for preventing infections, comprises administering an antibody including an engineered constant region.
   CTTGGGTGGACCATCCGTCTTCATCTTCCCTCCAAAGATCAAGGATGTACTCATGATCTC
  CTTGGGTGGACCATCCGTCTTCATCTTCCTCCAAAGATCAAGGATGTACTCATGATCTC
   786 TGAGCCCAGAGGGCCCACAATCAAGCCCTGTCCTCCATGCAAATGCCCAGCACCTAACCT
  Gaps
   / Match 47.2%; Score 700.6; DB 14; Length 1569; Local Similarity 100.0%; Pred. No. 2.8e-178; les 700; Conservative 0; Mismatches 0; Indels 0;
   Sequence 1569 BP; 446 A; 443 C; 356 G; 324 T; 0 U; 0 Other;
  Evans MJ;
  Squinto SP,
   Disclosure, SEQ ID NO 18; 87pp; English.
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D,
  30-MAY-2003; 2003US-0475202P.
  Rother RP, Faas-Knight S,
   (ALEX-) ALEXION PHARM INC.
   WPI; 2005-112860/12.
P-PSDB; ADW71833.
   846
   804
  906
  Query Match
  996
   Matches
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Pred. No. is the number of results predicted by chance to have a score greater than or equal to the score of the result being printed, and is derived by analysis of the total score distribution.

|           | Description              | Aaf55099 DNA encod | Aat04262 Hybrid IA | Adq31225 I-Ab(beta | Aat04269 Hybrid IA | Adq31228 I-Ab(beta | Aat86989 SCE1 sing | Aca60744 Mouse MHC | Aat86987 SSC1 sing | Aca60742 Mouse MHC | Aat86988 SCT1 sing | Aax89069 Single ch | Aca60743 Mouse MHC | Aat17588 Vector SC | Aat17586 Vector SS | Aat17587 Vector SC | Aav12068 Murine IA | Adj75986 Marker ge | Adx26090 Novel cel |
|-----------|--------------------------|--------------------|--------------------|--------------------|--------------------|--------------------|--------------------|--------------------|--------------------|--------------------|--------------------|--------------------|--------------------|--------------------|--------------------|--------------------|--------------------|--------------------|--------------------|
| SUMMARIES | αī                       | AAF55099           | AAT04262           | 2 ADQ31225         | AAT04269           | 2 ADQ31228         | AAT86989           | ACA60744           | AAT86987           | ACA60742           | AAT86988           | AAX89069           | ACA60743           | AAT17588           | AAT17586           | AAT17587           | AAV12068           | 2 ADJ75986         | 4 ADX26090         |
|           | Query<br>Match Length DB | 921 5              | 893 2              | 945 12             | 1013 2             | 915 12             | 1382 2             | 1382 8             | 1385 2             | 1385 8             | 1508 2             | 1508 2             | 1508 8             | 1382 2             | 1385 2             | 1508 2             | 4724 2             | 798 12             | 798 14             |
| عِن       | Query<br>Match I         | 96.8               | 74.6               | 69.9               | 68.1               | 66.1               | 63.9               | 63.9               | 63.9               | 63.9               | 63.9               | 63.9               | 63.9               | 63.7               | 63.7               | 63.7               | 62.9               | 59.1               | 59.1               |
|           | Score                    | 921                | 709.2              | 665.2              | 648                | 628.6              | 607.8              | 607.8              |                    |                    | 607.8              |                    | 607.8              | 606.2              | 606.2              | 606.2              | 598.6              | 561.8              | 561.8              |
|           | ę i                      | п                  | ~                  | ო                  | 4                  | Ŋ                  | y                  | ,                  | ∞                  | o                  | 2                  | 11                 | 12                 | 13                 | 14                 | 15                 | 16                 | 11                 | 18                 |

Example 1; Page 34-35; 43pp; French.

| Ab199040 Murine pC |            | Aaq56920 Mouse I-A | Aag35055 IAB beta | _          | Abn84048 Single ch | Abi99039 Murine pC | _          | Abi99028 IAS MBP 1 |            | Abi99027 IAS MBP 1 | Abi99021 I-As MBP. | IAS        | IAS        |            | Abk63510 Rat seque | Adb57995 Toxicity- |             | Adw21868 Rat hepat |             | Adx25826 Novel cel | Adg38634 Human SNP | Adq38637 Human SNP | Abk84087 Human cDN | Abx63009 Human cDN | Aad63150 Human maj |  |
|--------------------|------------|--------------------|-------------------|------------|--------------------|--------------------|------------|--------------------|------------|--------------------|--------------------|------------|------------|------------|--------------------|--------------------|-------------|--------------------|-------------|--------------------|--------------------|--------------------|--------------------|--------------------|--------------------|--|
| 4 ABI99040         | 2 AAT06286 | 2 AAQ56920         | 2 AAQ35055        | 4 ABI99038 | 6 ABN84048         | 4 ABI99039         | 4 ABI99031 | 4 ABI99028         | 4 ABI99032 | 4 ABI99027         | 4 ABI99021         | 4 ABI99030 | 4 ABI99029 | 4 ABI99033 | 6 ABK63510         | 10 ADB57995        | 10 ABT41775 | 11 ADW21868        | 13 ADV40851 | 14 ADX25826        | 13 ADQ38634        | 13 ADQ38637        | 6 ABK84087         | 8 ABX63009         | 10 AAD63150        |  |
| 1085               | 707        | 702                | 702               | 1698       | 1243               | 1662               | 1686       | 1701               | 2059       | 2346               | 1680               | 1707       | 2053       | 2343       | 562                | 299                | 562         | 562                | 562         | 562                | 1869               | 1892               | 1111               | 1199               | 1192               |  |
| 57.8               | 57.0       | 57.0               | 56.3              | 55.2       | 52.8               | 52.5               | 52.3       | 52.3               | 52.3       | 52.3               | 52.2               | 52.2       | 52.2       | 52.2       | 45.6               | 45.6               | 45.6        | 45.6               | 45.6        | 45.6               | 43.6               | 43.6               | 43.2               | 43.2               | 42.9               |  |
| 549.8              | 542        | 542                | 535.6             | 525.4      | 502.6              | 499.4              | 497.2      | 497.2              | 497.2      | 497.2              | 496                | 496        | 496        | 496        | 433.6              | 433.6              | 433.6       | 433.6              | 433.6       | 433.6              | 414.8              | 414.8              | 410.8              | 410.8              | 407.6              |  |
| 19                 | 2 2        | 22                 | 23                | 24         | 25                 | 26                 | 27         | 28                 | 29         | 30                 | 31                 | 32         | 33         | 34         | 35                 | 36                 | 37          | 38                 | 39          | 40                 | 41                 | 42                 | 43                 | 44                 | 45                 |  |

## ALIGNMENTS

AAF55099 standard; DNA; 921 BP.

AAF55099;

New soluble recombinant protein, useful e.g. as immunostimulant, comprises dimeric major histocompatibility complex molecule fused to immunoglobulin Fc region. Recombinant protein; alpha chain; beta chain; MHC; immunoglobulin; major histocompatibility complex; Fc region; antigen; T lymphocyte; immunostimulant; vaccine; infection; tumour; 88. DNA encoding a fusion protein comprising a beta chain of MHC. Location/Qualifiers (CNRS ) CNRS CENT NAT RECH SCI. 29-JUL-1999; 99FR-00009862. 28-JUL-2000; 2000WO-FR002193. Glaichenhaus N, Malherbe L; 15-MAY-2001 (first entry) WPI; 2001-182944/18. P-PSDB; AAB67481. WO200109194-A1 08-FEB-2001. Synthetic. ARESULT 1
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The specification describes soluble recombinant proteins that comprise at least a dimer formed from the alpha and beta-chains of MHC (major histocompatibility complex) Class I and II molecules in which at least one chain has, attached to its C-terminus, at least part of the Fc region of an immunoglobulin. The recombinant proteins, when linked to an antigening peptide, are used to count and/or purity antigen-reactive T lymphocytes and to characterize their phenotype, e.g. in preclinical evaluation of vaccines. They are also used as immunostimulants, particularly for vaccine development (against infections and tumours), to count and determine phenotype of autoreactive T cells in subjects with, or at risk of developing, autoimmune diseases, e.g. for staging or extisk of developing, autoimmune diseases, e.g. for staging or evaluating treatments, and (to purify and/or enrich Ag-reactive T cells from cell cultures or preventative cellular therapy. The present sequence encodes a recombinant protein of the invention, comprising a beta chain of MHC molecules

Sequence 921 BP; 214 A; 265 C; 286 G; 156 T; 0 U; 0 Other;

ô 140 720 240 320 440 420 480 560 540 620 600 680 99 740 800 80 9 GGGACGCAGCGCATACGGCTCGTGACCAGATACATCTACAACCAGGAGGAGGAGTACGTGCGC GTCATGCTGGAGATGACCCCTCATCAGGAGAGGTCTACACCTGCCATGTGGAGCATCCC GGTGGAGGCTCCGAAAGGCATTTCGTGGTTCCAGTTCAAGGGCGAGTGCTACTACACCAAC 361 TACTGGAACAGCCAGCCGGAGATCCTGGAGCGAACGCGGGCCGAGGTGGACACGCGGCGGC AGACACAACTACGAGGGGCCGGAGACCAGCACCTCCCTGCGGCGGCTTGAACAGCCCAAT GTGGCCATCTCCCTGTCCAGGACAGAGGCCCTCAACCACCACAAACACTCTGGTCTGTTCG GTGACAGATTTCTACCCAGCCAAGATCAAAGTGCGCTGGTTCAGGAATGGCCAGGAGGAG GTGACAGATTTCTACCCAGCCAAGATCAAAGTGCGCTGGTTCAGGAATGGCCAGGAGGAG ACAGTGGGGGTCTCATCCACAGCTTATTAGGAATGGGGACTGGACCTTCCAGGTCCTG 21 ATGGCTCTGCAGATCCCCAGCCTCCTCTCAGCTGCTGTGGTGGTGTTGATGGTGCTG AGCAGCCCCGGGACTGAGGGCGGAAACTCCATCTGCTTCTCGCCGTCGCTGGAGCACCCG ATCGTGGTGTGGGACGGAGGTGGGGGCTCACTAGTGCCCCGAGGCTCTGGA GGGACGCAGGCGCATACGGCTCGTGACCAGATACATCTACAACCGGGAGGAGTACGTGCGC TACGACAGGACGTGGCGAGTACCGCGGTGACCGAGCTGGGGCGGCCAGACGCCGAG 301 raceacadedacereseceastraceseceseseaseceasereseceseceseceases TACTIGGAACAGCCAGCCGGAGATCCTGGAGCGAACGCGGGGCCGAGGTGGACACGGCGTGC AGACACAACTACGAGGGCCGGAGACCAGCACCTCCCTGCGGCGCTTGAACAGCCCAAT GTCGCCATCTCCCTGTCCAGGACAGAGGCCCTCAACCACCACAACACTCTGGTCTGTTCG ACAGTGGGGGTCTCATCCACACACCTTATTAGGAATGGGGACTGGACCTTCCAGGTCCTG GTCATGCTGGAGATGACCCCTCATCAGGAGGGTCTACACCTGCCATGTGGAGCATCCC AGCCTGAAGAGCCCCATCACTGTGGAGTGGAGCGCACAGTCCGAGTCTGCCCGGAGCAAG 0; Gaps Length 921; 0, Indels ery Match 96.8%; Score 921; DB 5; Le st Local Similarity 100.0%; Pred. No. 8.4e-205; tches 921; Conservative 0; Mismatches 0; 121 421 621 601 741 81 61 141 201 181 261 241 321 381 441 501 481 561 541 681 661

WO9523814-A1

920 GGAGGTGGAGGATCCACTACAGCTCCATCAGCTCAGTTGAAAAAGAAATTGCAAGCACTG /trag= d /note= leader region" /note= leader region" /trag= e /note= "binding site for primer #331 (see AAT04261)" /trag= f \*tag= b note= "binding site for primer #261 (see AAT04260)" /\*tag= m /note= "probable primer binding site (primer #232)" 371. .389 /\*teg= h /note= "probable primer binding site (primer #270)" /Inte= "probable primer binding site (primer #270)" note= "probable primer binding site (primer #271)" /\*tag= k /note= "probable primer binding site (primer #272)" 808. .836 note= "probable primer binding site (primer #259)" 177. .893 1. 16
 /\*tag= a
 /note= "probable primer binding site (primer #233)"
 complement (45. 74) \*tag= g note= "primer #333 (see AAT04263) binding site" Polymerase chain reaction; PCR; primer; amplify; major histocompatability complex; MHC; T-cell receptor; TCR; autoimmune disease; immunodeficiency disease; immune response; immunoproliferation disease; graft-host rejection; therapy; 88. "IA beta chain beta 2 region" /note= "primer #332 binding site" 61..828 /\*tag= c /product= hybrid IA beta chain" complement (199. .250) Location/Qualifiers CATCATCATCATCATTGA 941 CATCATCATCATCATTGA 921 BP AAT04262 standard; DNA; 893 /product= 521. .550 /\*tag= j (first entry) Hybrid IA beta chain gene. \*tag= misc feature 16-APR-1996 Key primer\_bind primer\_bind mat\_peptide primer\_bind sig\_peptide primer\_bind primer\_bind primer bind primer\_bind primer\_bind primer\_bind 841 Synthetic AAT04262; 921 901 861 901 RESULT 2 AAT04262 g 셤 g ઠે ò ò

607

667 692 727 752 787

ed Jun 28

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Peptide-Class II major histocompatibility complex (MHC) composite, useful for detecting antigen specific CD4+ T-cell, comprises antigen peptide containing epitope of mucous membrane invasive protein, and extracellular region of MHC.
  The invention relates to a novel class II major histocompatibility complex (MHC) antigenic peptide composite comprising a peptide containing the T-cell antigenic determinant of a mucous membrane invasive protein and the extracellular region of a class II MHC molecule or at least part of the extracellular region of the class II MHC molecule having an amino
   I-Ab(beta)-Cholera toxin B subunit-leucine zipper (LZ)-BirA fusion cDNA.
   class II major histocompatibility complex; WHC; CD4+ T-cell detection; flow cytometry; mucous membrane invasive antigen; I-Ab(beta)-cholera toxin B subunit-leucine zipper-BirA fusion; CTB; 88;
   693 CTTCCAGGTCCTGGTCATGCTGGAGATGACCCCTCATCAGGGAGAGGGTCTACACCTGCCA
   TGTGGAGCATCCCAGCCTGAAGAGCCCCCATCACTGTGGAGTGGAGGGCCACAGTCCGAGTC
  TCTGGTCTGGTTCGGTGACAGATTTCTACCCAGCCAAGATCAAAGTGCGCTGGTTCAGGAA
  573 reregrenerregereachearriceaceaceachacheangreceregereach
   TGGCCAGGAGGAGACAGTGGGGGTCTCATCCACACAGCTTATTAGGAATGGGGACTGGAC
  CTTCCAGGTCCTGGTCATGCTGGAGATGACCCCTCATCAGGGAGAGGGTCTACACCTGCCA
   1. .945
/*tag= a
/product= "I-Ab(beta)-Cholera toxin B subunit (CTB)-
leucine zipper (LZ)-BirA fusion cDNA"
  (SENT-) SENTAN KAGAKU GLJUTSU INCUBATION CENT KK.
   Example 1; SEQ ID NO 10; 30pp; Japanese.
   TGCCCGGAGCAAGGGAGGTGGAGG 811
  Location/Qualifiers
  BP.
  ADQ31225 standard; cDNA; 945
   03-DEC-2003; 2003JP-00404367.
   03-DEC-2002; 2002JP-00351818.
   07-OCT-2004 (first entry)
  WPI; 2004-546819/53.
P-PSDB; ADQ31224.
  Vibrio cholerae.
Unidentified.
  JP2004196789-A.
  15-JUL-2004.
  548
   609
  668
  ADQ31225;
                            513
   728
  788
   gene.
   RESULT 3
ADQ31225
ID ADQ3
  셤
   셤
   ઠે
   셤
   ጵ
  ઠે
  g
   ઠે
  ઠે
   This sequence represents a hybrid IA beta chain gene, containing the chicken ovalbumin peptide (COVA). This sequence was used in the construction of a hybrid IA alpha beta dimer. The encoded protein (plAd-OVA) was found to be more stable than the IA alpha beta dimer. The stability was decreased by the addition of a MHC groove specific binding peptide (e.g. see AAR82527, AAR82528 and AAR82531), compared to an increase seen on the addition of a MHC binding peptide to IE k/d-MCC. These complexes may be used to regulate an immune response. The complexes are capable of being recognised by a TCR alone or in combination with additional MHC proteins. These complexes are useful for therapeutic purposes and experimental purposes. They can also be used as reagents for the treatment of diseases including autoimmune diseases, immunodeficiency
   7
   162
   212
   247
   272
   367
  392
   427
  452
  487
  48 GGAATTCTTAGAGATGGCTCTGCAGATCCCCAGCCTCCTCTCAGCTGCTGTGTGGT 107
   68 GCTGATGGTGCTGAGCAGCCCCGGGACTGAGGCGCGGAAACTCCATCTGCTTCTCGCCGTC 127
  128 GCTGGAGCACCCGATCGTGGTGTCCGGCAGCTGGGACGAGGTGGGGGCTCACTAGTGCC 187
  CTACTACACCAACGGGACGCGCATACGGCTCGTGACCAGATACATCTACAACCGGGA 307
  332
  488 TGAACAGCCCAATGTCGCCATCTCCCTGTCCAGGACAGAGGCCCTCAACCACCACAACAC 547
  Peptide-MHC complex comprising antigenic peptide, linker and MHC segment - useful as reagents for the treatment of diseases including auto-immune diseases, immuno-stimulatory diseases or graft-host rejection.
  67
   108 GCTGATGGTGCTGAGCAGCCCCGGGAACTGAGGGCGGAAACTCC----GTACATGCTGCC
  163 CATGCTGAGATCAATGAGGCTGGCAG-----AGGAGGTGGGGGGTCACTAGTGCC
   CCGAGGCTCTGGAGGTGGAGGCTCCGAAAGGCATTTCGTGGTCCAGTTCAAGGGCGAGTG
   CCGAGGCTCTGGAGGTGGAGGCTCCGAAAGGCATTTCGTGGTCCAGTTCAAGGGCGAGTG
   GGAGTACGTGCGCTACGACGACGTGGGCGAGTACCGCGCGCTGACCGAGCTGGGGCG
  333 GGAGTACGTGCGCTACGACAGCGACGTGGGCGAGTACCGCGCGGTGACCGAGCTGGGGCG
  453 GGACACGGCGTGCAGACACAACTACGAGGGCCGGAGACCAGCACCTCCCTGCGGCGCTT
   CTACTACACCAACGGGACGCAGGGCATACGGCTCGTGACCAGATACATCTACAACCGGGA
   393 GCCAGACGCCGAGTACTGGAACAGCCAGCCGGAGATCCTGGAGCGAACGCGGGCCGAGGT
  8 GGAATTCTTAGAGATGGCTCTGCAGATCCCCAGCCTCCTCCTCTCAGCTGCTGGTGGT
   GCCAGACGCCGAGTACTGGAACAGCCAGCCGGAGATCCTGGAGCGAACGCGGGCCCGAGGT
   Gaps
   diseases, immunoproliferation diseases, and graft-host rejection
   Indels 15;
  Length 893;
  Sequence 893 BP; 204 A; 239 C; 275 G; 175 T; 0 U; 0 Other;
  lery Match
74.6%; Score 709.2; DB 2;
set Local Similarity 94.7%; Pred. No. 2.1e-155;
stches 761; Conservative 0; Mismatches 28;
   (NAJE-) NAT JEWISH CENT IMMUNOLOGY & RESPIRATORY
  Example 1; Page 53; 94pp; English.
  95WO-US002689
   94US-00207481
  Marrack P;
  WPI; 1995-320543/41.
P-PSDB; AAR82533.
  03-MAR-1995;
   04-MAR-1994;
  Kappler JW,
                              08-SEP-1995
   188
  213
  248
   273
   368
  428
   308
```

```
acid sequence comprising one or more deletions, substitutions or additions. The molecule of the invention may be useful for detecting an antigen-appecific CD4+ T-cell by flow cytometry and for presenting a microorganism-derived mucous membrane invasive protein as an antigen. The method of the invention enables efficient detection of antigen-specific activation of CD4+ T-cells in the mucous membrane. The current sequence is that of the class II major histocompatibility complex-related I-Ab(alpha)-Cholera toxin B subunit (CTB)-leucine zipper (LZ)-BirA fusion cDNA of the invention.
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Seguence 945 BP; 230 A; 256 C; 294 G; 165 T; 0 U; 0 Other;

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ñ
   182
  225
  302
   362
  345
  422
  405
   482
   465
   645
  705
   782
   765
  902
   879
  108
   242
  542
   525
  602
   585
   662
  722
  842
  819
  123 CCGTCGCTGGAGCACCCGATCGTGGTGTCCGGCAGCTGGGACGGAGGTGGGGGCTCACTA
   AATAAGACGCCGCACGCGATCGCGGCCATCAGCATGGCGAACGGAGGTGGTGGTCGGT
   GGGCGGCCAGACGCCGAGTACTGGAACAGCCAGCCGGAGATCCTGGAGCGAACGCGGGCC
  CGGCTTGAACAGCCCAATGTCGCCATCTCCCTGTCCAGGACAGAGGCCCTCAACCACCAC
   526 AACACTCTGGTCTGCTCAGTGACAGATTCTACCCAGCCAAGATCAAAGTGCGCTGGTTC
   AGGAATGGCCAGGAGGAGACAGTGGGGGTCTCATCCACACAGCTTATTAGGAATGGGGAC
   CGGAATGGCCAGGAGGAGACGGTGGGGGTCTCATCCACACAGCTTATTAGGAATGGGGAC
  TGGACCTTCCAGGTCCTGGTCATGCTGGAGATGACCCCTCATCAGGGAGAGGTCTACACC
   TGTCACGTGGAGCATCCCAGCCTGAAGAGCCCCATCACTGTGGAGTGGAGGGCACAGTCG
   AAGAAATTGCAAGGCACTGAAGAAAAAAAGACGCTCAGCTGAAGTGGAAACTTCAAGCCCTC
  GTGGTGCTGATGCTGAGCAGCCCCGGGACTGAGGGCGGAAACTCCATCTGCTTCTCG
  GTGCCCCGAGGCTCTGGAGGTGGAGGCTCCGAAAGGCATTTCGTGGTCCAGTTCAAGGCC
  GAGTGCTACTACACCAACGGGACGCACATACGGCTCGTGACCAGATACATCTACAAC
   GAGTGCTACTTCACCAACGCGACGCACACACATACGATATGTGACCAGATACATCTACAAC
   GGGCGGCCAGACGCCGAGTACTGGAACAGCCGGAGATCCTGGAGCGAACGCGGGCC
   GAGGTGGACACGCCTCCACACACAACTACGAGGGCCCGGAGACCAGCACCTCCCTGCGG
  543 AACACTCTGGTCTGGTTCGGTGACAGATTTCTACCCAGCCAAGATCAAAGTGCGCTGGTTC
   646 redaccrrccaegrecregarcarecregaeareaceceregeegeagaegreracaee
   TGCCATGTGGAGCATCCCAGCCTGAAGAGCCCCATCACTGTGGAGTGGAGGGCACAGTCC
  GAGTCTGCCCGGAGCAAGGGAGGTGGAGGATCCACTACAGCTCCATCAGCTCAGTTGAAA
                                    Gaps
                                  6
    Length 945;
/ Match 69.9%; Score 665.2; DB 12; Length Local Similarity 86.6%; Pred. No. 3.8e-145; nes 759; Conservative 0; Mismatches 108; Indels
   903 AAGAAGAAACTCGCCCAGCATCATCATCATCAT 938
  109
   406
  483
  603
   63
   49
  466
  183
   169
   243
   226
  303
   363
   346
   423
  586
   663
  723
   206
  783
   843
    ery Match
                     at Loca
tches
```

/\*tag= m //tag= m //tag= morphable primer binding site (primer #366)" //tag= n //tag= n //tag= n //tag= n //tag= n //tag= n //tag= mprobable primer binding site (primer #59)"

94US-00207481

04-MAR-1994;

WO9523814-A1

08-SEP-1995

primer\_bind

primer\_bind

```
/*tag= e
/note= "binding site for primer #363 (see AAT04271)"
complement(177. .226)
  /*tag= b
/note= "binding site for primer #362 (see AAT04270)"
  385. 403
/*tag= h
//nce= "probable primer binding site (primer #270)"
//tag= 1
   /noce= "probable primer binding site (primer #271)"
544. .568
/*tag= k
/*tag= k
923. .850
  /*tag= 1
/note= "probable primer binding site (primer #259)"
942. .976
  Location/Qualifiers
1. .18
//tote= "probable primer binding site (primer #76)"
complement(40. .74)
   Major histocompatability complex; MHC; T-cell receptor; TCR; autoimmune disease; immunodeficiency disease; immune response; immunoproliferation disease; graft-host rejection; therapy; B M12.C3; pM12-IAb-Ea; ss.
   /product= "IA beta chain beta 2 region"
535. .564
   /*tag= c /*tag= c /*tag= c /*tag= c /*tag= d /*tag= d /*note= "leader region"
          915
  /*tag= f
/note= "primer #364 binding site"
complement(212. .266)
BP.
   standard; DNA; 1013
  (first entry)
   Hybrid IA beta chain gene
  *tag=
   16-APR-1996
  primer_bind
   primer_bind
  primer_bind
   sig_peptide
   primer_bind
   primer_bind
  primer_bind
   primer_bind
  peptide
  primer_bind
  primer_bind
   Synthetic
   AAT04269
           880
   AAT04269
  RESULT
           셤
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This sequence represents a hybrid IA beta chain gene. This sequence contains a fragment of the IB alpha chain (residues 56-73), as well as a linker and cleavage site. This sequence was transfected into a B cell line (M12.03) using plasmid pM12-IAb-EB. It was found that the encoded sequence was expressed in these cells. Complexes such as this may be used to regulate an immune response. The complexes are capable of being recognised by a TCR alone or in combination with additional MiC proteins. These complexes are useful for therapeutic purposes and experimental purposes. They can also be used as reseents for the treatment of diseases including autoimmune diseases, immunodeficiency diseases,
  67 TGCTGATGGTGCTGAGCAGCCCCGGGACTGAGGCGGGAAACTCCATCTGCTTCTCGCCGT 126
   127 CGCTGGAGCACCCCGATCGTGGTGTCCCGCCAGCTGGGACGTGGGGGGCTCACTAGTGC 186
   Peptide-MHC complex comprising antigenic peptide, linker and MHC segment - useful as reagents for the treatment of diseases including auto-immune diseases, immuno-stimulatory diseases or graft-host rejection.
   406 GGCCAGACGCCGAGTACTGGAACAGCCAGCCGGAGATCCTGGAGCGAACGCGGGCCGAGG
   7 GGGAATTCTTAGAGATGGCTCTGCAGATCCCCAGCCTCCTCTCTCAGCTGCTGTGGTGG
   169 crcaccerceacracarrecrereacaacaacarrecreacaacarecreacare---cegre
  226 GAGGGGGAAGTGGAGGTGTGAAAGGCATTTCGTGTACCAGTTCATGGGGAGT
  367 GGCCAGACGCCGAGTACTGGAACAGCCAGCCGGAGATCCTGGAGCGAACGCGGGCCGAGG
  TGGACACGCCGTGCAGACACAACTACGAGGGCCCGGAGACCAGCACCTCCCTGCGGCGGC
  466 TGGACACGGTGCAGACACAACTACGAGGGCCGGAGACCCACACCTCCCTGCGGCGGC
   247 GCTACTACACCAACGGGACGCAGCGCATACGGCTCGTGACCAGATACATCTACAACCGGG
  187 CCCGAGGCTCTGGAGGTGGAGGCTCCGAAAGGCATTTCGTGGTCCAGTTCAAGGGCGAGT
   Gaps
   3;
  68.1%; Score 648; DB 2; Length 1013; 89.3%; Pred. No. 4e-141; ive 1; Mismatches 81; Indels
  Sequence 1013 BP; 220 A; 272 C; 327 G; 192 T; 0 U; 2 Other;
(NAJE-) NAT JEWISH CENT IMMUNOLOGY & RESPIRATORY
  Example 2; Page 65; 94pp; English.
   set Local Similarity 89.3 stches 709, Conservative
   Marrack P;
   WPI; 1995-320543/41.
P-PSDB; AAR82538.
   Kappler JW,
  427
  ery Match
```

```
The invention relates to a novel class II major histocompatibility complex (MHC) antigenic peptide composite comprising a peptide containing the T-cell antigenic determinant of a mucous membrane invasive protein and the extracellular region of a class II MHC molecule or at least part of the extracellular region of the class II MHC molecule having an amino acid sequence comprising one or more deletions, substitutions or additions. The molecule of the invention may be useful for detecting an antigen-specific CD4+ T-cell by flow cytometry and for presenting a microcorganism derived mucous membrane invasive protein as an antigen. The method of the invention enables efficient detection of antigen-specific activation of CD4+ T-cells in the mucous membrane. The current sequence is that of the class II major histocompatibility complex-related I-bb (alpha)-Escherichia coli heat-labile toxin B subunit (ITB)-leucine zipper (IZ)-BirA fusion CDNA of the invention.
  Peptide-Class II major histocompatibility complex (MHC) composite, useful for detecting antigen specific CD4+ T-cell, comprises antigen peptide containing epitope of mucous membrane invasive protein, and extracellular region of MHC.
                              726
   825
646 ATGGCCAGGAGGAGACGGTGGGGCTCTCATCCACACACTGTTAGGAATGGGACTGGA 705
   /product= "I-Ab(beta)-Escherichia coli heat-labile toxin
B subunit (LTB)-leucine zipper (LZ)-BirA fusion protein"
   class II major histocompatibility complex; MHC; CD4+ T-cell detection; flow cytometry; mucous membrane invasive antigen; I-Ab(beta)-heat-labile toxin B subunit-leucine zipper-BirA fusion; LTB;
                                CCTTCCAGGTCCTGGTCATGCTGGAGATGACCCCTCATCAGGGAGAGGTCTACACCTGCC
  706 CCTTCCAGGTCCTGGTCATGCTGGAGATGACCCCTCGGCGGGGAGAGGTCTAYACCTGTC
   ACGIGGAGCATCCCAGCCTGAAGAGCCCCCATCACTGTGGAGGGGGGGCACAGTCTGAGT
   727 ATGREGAGCATCCCAGCCTGAAGAGCCCCCATCACTGTGGAGTGGAGGCCACAGTCCGAGT
  I-Ab(beta)-E. coli heat-labile toxin B subunit-LZ-BirA fusion cDNA.
   (SENT-) SENTAN KAGAKU GLJUTSU INCUBATION CENT KK.
  Example 3; SEQ ID NO 13; 30pp; Japanese.
  Location/Qualifiers
   ADQ31228 standard; cDNA; 915 BP.
  03-DEC-2003; 2003JP-00404367
  03-DEC-2002; 2002JP-00351818
  826 criccriccaccase 839
   CTGCCCGGAGCAAG 800
  (first entry)
  /*tag= a
  1. .915
   WPI; 2004-546819/53.
P-PSDB; ADQ31227.
   Escherichia coli.
Unidentified.
   JP2004196789-A.
  07-0CT-2004
  15-JUL-2004.
  787
   ADQ31228;
                                667
   ss; gene.
   994
  RESULT 5
   ADQ31228
   요
   ð
   g
  ò
  g
                              ઠે
  108
   168
   225
  246
  285
   306
   345
   366
  405
  426
  465
   486
   525
   546
  585
  909
   645
   999
   99
   526 TTGAACAGCCCAATGTCGTCATCTCCCTGTCCAGGACAGAGGCCCTCAACACAACA
  ATGGCCAGAGAGAGACAGTGGGGGTCTCATCCACAGCTTATTAGGAATGGGGACTGGA
  CTCTGGTCTGTTCGGTGACAGATTTCTACCCAGCCAAGATCAAAGTGCGCTGGTTCAGGA
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요
  1;
   748
  808
  868
  261
  921
  214
   321
   274
  381
  441
   394
  561
  514
   621
   574
   681
   634
  741
  694
   GCCTGAAGAGCCCCATCACTGTGGAGTGGAGGGCACAGTCCGAGTCTGCCCGGAGCAAGG 801
   GAGGIGGAGGAICCACIACAGCICCAICAGCICAGTIGAAAAAGAAATIGCAAGCACIGA 861
   501
   454
   ACTGGAACAGCCAGCCGGAGATCCTGGAGCGAACGCGGGCCGAGCTGGACACGGTGTGCA
   155 GAGGTGGTAGTGAAAGGCATTTCGTGTACCAGTTCATGGGCGAGTGCTACTTCACCAACG
  ACGACAGCGACGTGGGCGAGTACCGCGCGGGGGCGAGCTGGGGCGGCCGAGACGCCGAGT
   GACACAACTACGAGGGGCCGGAGACCCACCTCCCTGCGGGGGGCGCTTGAACAGCCCAATG
  TCGTCATCTCCCTGTCCAGGACAGACCCTCAACCACCACAAAACTTCTGGTCTGCTCAG
   CAGTGGGGGTCTCATCCACACACATATTAGGAATGGGGACTGGACCTTCCAGGTCCTGG
   cecrecescretecacacacerrarrageaareeseacreceacereces
  AGAAAAAGAACGCTCAGCTGAAGTGGAAACTTCAAGCCTCAAGAAGAAACTCGCCCAGC
   202 GIGGAGGCICCGAAAGGCAITICGIGGICCAGIICAAGGGCGAGIGCIACTACAACG
   GGACGCAGCGCATACGGCTCGTGACCAGATACATCTACAACCGGGAGGAGTACGTGCGCT
   215 GGACGCAGCGCATACGATATGTGACCAGATACATCTACAACCGGGAGGAGTACGTGCGCT
  ACTEGAACAGCCAGCCGGAGATCCTGGAGCGAACGCGGGCCGAGGTGGACACGCGTGCA
   GACACAACTACGAGGGCCGGAGACCAGCACCTCCCTGCGGCGGCTTGAACAGCCCAATG
  TCGCCATCTCCCTGTCCAGGACAGAGGCCCTCAACCACCACAACACTCTGGTTCGG
   TGACAGATTTCTACCCAGCCAAGATCAAAGTGCGCTGGTTCAGGAATGGCCAGGAGGA
   TGACAGATTTCTACCCAGCCAAGATCAAAGTGCGCTGGTTCCGGAATGGCCAGGAGGAGA
  TCATGCTGGAGATGACCCCTCATCAGGGAGAGGTCTACACCTGCCATGTGGAGCATCCCA
  Gaps
   Construction, major histocompatibility complex, MHC, fusion complex, SCE1 single chain gene, 88.
  9
                                     Length 915;
A; 242 C; 271 G; 174 T; 0 U; 0 Other;
                               ttch 66.1%; Score 628.6; DB 12; Length cal Similarity 91.9%; Pred. No. 1.3e-136; 677; Conservative 0; Mismatches 54; Indels
  BP
   922 ATCATCATCATCAT 938
  AAT86989 standard; DNA; 1382
   recarcararreregar
   (first entry)
   SCE1 single chain gene
 Sequence 915 BP; 228
   27-MAR-1998
   395
  635
  682
                                   ery Match
st Local S
tches 677
   262
  322
  275
  382
   335
   442
  502
  455
   562
   515
   622
  575
   742
   695
  802
  749
  862
  809
   869
   AAT86989;
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Single chain major histocompatibility complex comprising linked alpha and beta chains - useful for suppressing an immune response to an auto:Immune disease, e.g. multiple sclerosis, rheumatoid arthritis, diabetes mellitus, etc.
  81 AGCAGCCCGGGAACTGAGGGGGGGAAACTCCATCTGCTTCTCGCCGTCGCTGGAGCACCCG 140
   GGTGGAGGCTCCGAAAGGCATTTCGTGGTCCAGTTCAAGGGCGAGTGCTACTACACCAAC 260
  GTCGCCATCTCCCTGTCCAGGACAGAGGCCCTCAACCACCACAACACTCTGGTCTGTTCG 560
   105 ccrecrcacecreanicanceanderestrorecracesadeseseses 164
   GGGACGCAGCGCATACGGCTCGTGACCAGATACATCTACAACCGGGAGGAGTACGTGCGC 320
  141 ATCGTGGTGTCCGGCAGCTGGGAGGTGGGGCTCACTAGTGCCCCGAGGCTCTGGA
  TACGACAGCGACGTGGGCGAGTACCGCGCGGTGACCGAGCTGGGGCGCCCAGACGCCGAG
  AGACACAACTACGAGGGCCGGAGACCAGCACCTCCCTGCGGCGGCTTGAACAGCCCAAT
   TACTGGAACAGCCAGCCGGAGATCCTGGAGCGAACGCGGGCCGAGGTGGACACGGCGTGC
   21 ATGCTCTCCAGATCCCCAGCCTCCTCTCAGCTGCTGTGGTGGTGCTGATGGTGCTG
  21;
   Query Match 63.9%; Score 607.8; DB 2; Length 1382; Best Local Similarity 89.6%; Pred. No. 1e-131; Matches 673; Conservative 0; Mismatches 57; Indels 21;
  Sequence 1382 BP; 320 A; 373 C; 405 G; 284 T; 0 U; 0 Other;
  The present sequence was used in the construction of major histocompatibility complex (MHC) fusion complexes
  Wong HC;
  Example 17; Page 140-141; 217pp; English.
                   Location/Qualifiers
6. .1382
/*tag= a
  Rhode PR, Jiao J, Burkhardt M,
   97WO-US001617
  96US-00596387
  (DADE-) DADE INT INC
  WPI; 1997-402555/37.
   P-PSDB; AAW29214
  30-JAN-1997;
  31-JAN-1996;
   WO9728191-A1
   07-AUG-1997
Synthetic.
   261
   225
   285
   381
  441
   405
  465
  321
   201
   501
   Query Match
                       Key
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200

9

224

440

500

380

344

eg.

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584
  680
  644
  740
   704
GTGACAGATTTCTACCCAGGCCAAGATCAAAGTGCGCTGGTTCAGGAATGGCCAGGAGGAG
                         GTGACAGATTTCTACCCAGCCAAGATCAAAGTGCGCTGGTTCAGGAATGGCCAGGAGGAG
  ACAGTGGGGGTCTCATCCACACGTTATTAGGAATGGGGACTGGACCTTCCAGGTCCTG
  585 ACAGTGGGGGTCTCATCCACACACGCTTATTAGGAATGGGGGACTTGCAGGTCCTG
   645 GTCATGCTGGAGATGACCCCTCATCAGGGAGAGGTCTACACCTGCCATGTGGAGGATCCC
  GICATGCTGGAGATGACCCCTCATCAGGGAGGGTCTACACCTGCCATGTGGAGCATCCC
  AGCCTGAAGACCCCATCACTGTGGAGTGGA 735
   741 AGCCTGAAGAGCCCCATCACTGTGGAGTGGA
  621
  561
   525
  681
  705
```

BP ACA60744 standard; DNA; 1382

ACA60744;

16-JUN-2003 (first entry)

Mouse MHC I-Ad/Ova 323-339 synthetic gene SCE1

MHC; major histocompatibility complex; gene therapy; fusion complex; peptide-binding groove; T cell modulation; class II MHC; vaccine; autoimmune disorder; multiple sclerosis; rheumatoid arthritis; insulin-dependent diabetes mellitus; myasthenia gravis; immunogen; chronic allergy; mouse; ds; I-Ad; gene.

Synthetic.

US2002198144-A1.

26-DEC-2002

06-JUL-2001; 2001US-00900379.

94US-00283302. 95US-00382454. 97US-00776084. 29-JUL-1994; 01-FEB-1995; 17-JAN-1997;

DADE-) DADE INT INC.

Rhode PR, Weidanz JA, Grammer S, Edwards AC; Chavaillaz P, Jiao JJJ; Wong HC,

WPI; 2003-341126/32.

P-PSDB; ABU72108

Novel major histocompatibility complex fusion complex having presenting peptide covalently linked to MHC molecule containing peptide-binding groove, used for suppressing immune response in multiple sclerosis, allergies.

Example 17; Fig 29; 126pp; English.

The invention relates to a major histocompatibility complex (MHC) fusion complex (I) comprising an MHC molecule that contains a peptide-binding groove, and a presenting peptide covalently (e.g. an antigent peptide) linked to the MHC molecule, where (I) is capable of modulating the activity of a T cell. Also included are a DNA construct coding to the complex, where the MHC molecule is a class II MHC (e.g. mouse I-Ad or I-As, or human HLA-DRI (human leukocyte antigen-DRI), a multivalent MHC fusion complex comprising two or more linked complexes, identifying a peptide that can modulate the activity of T cells (involving introducing funch host cells aloning vectors that each contain the fusion complex DNA, culturing the host cells under conditable for expression of the MHC fusion complex, and selecting host cells that express MHC fusion

680

ACAGTGGGGGTCTCATCCACCACAGCTTATTAGGAATGGGGACTGGACCTTCCAGGTCCTG

621

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complex that modulate the activity of T cells), a single recombinant expression vector comprising DNA that codes for the alpha and beta chains of the fusion complex MHC protein, a single recombinant expression vector comprising DNA that codes for a T cell costimulatory factor and the alpha and beta chains of the MHC fusion complex. The DNA constructs can contain the expression of the fusion complex. Also included are inducing an immune cresponse in a mammal (including vaccinating a mammal against a targeted disorder, by administering DNA sequence comprising a fusion complex, or DNA sequence coding for a fusion complex which is a single chain fusion complex and subpressing an immune response in a mammal by administering to the mammal a DNA sequence comprising an expression vector, encoding a coding for a fusion complex which is a single chain fusion complex which in a series of main fusion complex which is a single chain fusion as transmembrane domain, and a presenting peptide that is a T cell receptor (TCR) antagonist or partial agonist and is covalently linked to the MHC protein, or DNA sequence complex which is a single chain fusion molecule. The methods are useful for identifying a peptide that can modulate the cativity of T cells, inducing an immune response in a mammal (including vaccinating a mammal inducing an include an autoimmune disorder cumuch as multiple sclerosis, insulin-dependent disorder; and abbetes mallitus, creative of the mammal and mammal. The disorder include an autoimmune disorder includes a mouse MHC class II I-Ad fusion complex of the ä 81 ACCAGCCCCGGGACTGAGGGCGGAAACTCCATCTGCTTCTCGCCGTCGCTGGAGCACCCG 140 165 GGGGGAAACTCCGAAAGGCATTTCGTGGTCCAGTTCAAGGGCGAGTGCTACACCAAC 224 320 344 440 AGACACAACTACGAGGGGCCGGAGACCAGCACCTCCCTGCGGCGGCTTGAACAGCCCCAAT 500 560 620 105 gengencacgengaancaacgaagenginegreeragegaggggagggaaggggggga 164 GGTGGAGGCTCCGAAAGGCATTTCGTGGTCCAGTTCAAGGGCGAGTGCTACTACACCAAC 260 284 321 TACGACAGCGACGAGGCGAGTACCGCGCGTGACCGAGCTGGGGCGGCCAGACGCCGAG 380 404 405 AGACACAACTACGAGGGGCCGGAGACCAGCACCTCCCTGCGGGCGCCGCTTGAACAGCCCCAAT 464 524 525 GTGACAGATTTCTACCCAGCCAAGATCAAAGTGCGCTGGTTCAGGAATGGCCAGGAGGAG 584 65 225 GGGACGCAGCGCATACGGCTCGTGACCAGATACATCTACAACCGGGAGGAGGAGTACGTGCGC 285 TACGACACGACGTGGGCGAGTACCGCGCGGGGACCTGGGGCGGCCAGACGCCGAG 345 TACTGGAACAGCCGGAGAGATCCTGGAGCGAACGCGGGCCGAGGTGGACACGGCGTGC GTCGCCATCTCCCTGTCCAGGACAGAGGCCCTCAACCACCACAAACACTCTGGTCTGTTCG 561 GTGACAGATTTCTACCCAGCCAAGATCAAAGTGCGCTGGTTCAGGAATGGCCAGGAGGAG GGGACGCAGCGCATACGGCTCGTGACCAGATACATCTACAACCGGGAGGAGTACGTGCGC TACTGGAACAGCCAGCCGGAGATCCTGGAGCGAACGCGGGCCGAGGTGGACACGGCGTGC 141 ATCGTGGTGTCCGGCAGCTGGGACGGAGGTGGGGGCTCACTAGTGCCCCCGAGGCTCTGGA Gaps Pred. No. 1e-131; 0; Mismatches 57; Indels 21; Score 607.8; DB 8; Length 1382; Sequence 1382 BP; 320 A; 373 C; 405 G; 284 T; 0 U; 0 Other; / Match 63.9%; Local Similarity 89.6%; les 673; Conservative ( invention 261 381 441 465 Query Match 201 501 Matches 유 셤 g 셤 유 ઠે 셤 ð ઠ 셤 ઠે ŝ 셤 ઠે δ 용 Š ò 셤 ò

224 320 284 344 440 560 524 620 680 644

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380
   404
   200
  464
  GTCATGCTGGAGATGACCCCTCATCAGGGAGAGGTCTACACCTGCCATGTGGAGCATCCC 740
   704
165 eccesaracticcerraciticeresticasticaresceresticares
   GTGACAGATTTCTACCCAGCCAAGATCAAAGTGCGCTGGTTCAGGAATGGCCAGGAGGAG
   GGGACGCAGCGCATACGGCTCGTGACCAGATACATCTACAACCGGGAGGAGTACGTGCGC
  225 GGGACGCACATACGGCTCGTGACCAGATACATCTACAACCGGGAGGAGTACGTGCGC
   TACGACAGCGACGTGGGCGAGTACCGCGCGGTGACCGAGCTGGGGCGGCCCAGACGCCGAG
  AGACACAACTACGAGGGCCGGAGACCAGCACCTCCCTGCGGCGCCTTGAACAGCCCAAT
  ACAGTGGGGGTCTCATCCACACACTTATTAGGAATGGGGACTGGACTTCCAGGTCCTG
   TACTGGAACAGCCAGCCGGAGATCCTGGAGCGAACGCGGGCCCGAGGTGGACACGCCGTGC
  MHC; major histocompatibility complex; gene therapy; fusion complex; peptide-binding groove; T cell modulation; class II MHC; vaccine; autochmune disorder; multiple sclerosis; rheumatoid arthritis; insulin-dependent diabetes mellitus; myasthenia gravis; immunogen; chronic allergy; mouse; ds; I-Ad; gene.
   Edwards AC;
   Mouse MHC I-Ad/Ova 323-339 synthetic gene SSC1
   Grammer S,
   AGCCTGAAGAGCCCCATCACTGTGGAGTGGA 771
   Rhode PR, Weidanz JA,
  BP
  94US-00283302.
95US-00382454.
97US-00776084.
  ACA60742 standard; DNA; 1385
  06-JUL-2001; 2001US-00900379
   16-JUN-2003 (first entry)
  (DADE-) DADE INT INC.
  US2002198144-A1
   29-JUL-1994;
01-FEB-1995;
17-JAN-1997;
  26-DEC-2002
  Synthetic
   441
   741
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  ACA60742;
   381
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   Single chain major histocompatibility complex comprising linked alpha and beta chains - useful for suppressing an immune response to an auto:immune disease, e.g. multiple sclerosis, rheumatoid arthritis, diabetes mellitus, etc.
   164
   81 AGCAGCCCCGGGACTGAGGGCGGAAACTCCATCTGCTTCTCGCCGTCGCTGGAGCACCCG 140
  141 ATCGTGGTGTCCGGCAGCTGGACGAGGTGGGGGCTCACTAGTGCCCCGAGGCTCTGGA 200
     644
   80
   65
  GTCATGCTGGAGATGACCCCTCATCAGGAGAGGTCTACACCTGCCATGTGGAGCATCCC
   6 ATGGCTCTGCAGATCCCCAGCCTCCTCTCAGCTGCTGTGGTGGTGCTGATGGTGCTG

    585 ACAGTGGGGGTCTCATCCACACACGTTATTAGGAATGGGGACTGGACCTTCCAGGTCCTG
                                GTCATGCTGGAGATGACCCCTCATCAGGGAGAGGTCTACACCTGCCATGTGGAGCATCCC
   21 ATGGCTCTGCAGATCCCCAGCCTCCTCTCAGCTGCTGTGGTGGTGCTGATGGTGCTG
   Gaps
   Construction, major histocompatibility complex, MHC, fusion complex, SSC1 single chain gene; 88.
   21;
   Length 1385;
   Sequence 1385 BP; 316 A; 383 C; 399 G; 287 T; 0 U; 0 Other;
   The present sequence was used in the construction of major histocompatibility complex (MHC) fusion complexes
   Indels
  Score 607.8; DB 2;
Pred. No. 1e-131;
0; Mismatches 57;
  741 AGCCTGAAGAGCCCCATCACTGTGGAGTGGA 771
  705 AGCCTGAAGAGCCCCATCACTGTGGAGTGGA 735
  HC;
  Example 17; Page 135-137; 217pp; English.
  Wong
   Location/Qualifiers
6. .1385
/*tag= a
  ΣÌ
  BP
  Burkhardt
  AAT86987 standard; DNA; 1385
  97WO-US001617
   96US-00596387
   ery Match 63.9%; st Local Similarity 89.6%; tches 673; Conservative
  (first entry)
   SSC1 single chain gene
  (DADE-) DADE INT INC.
   WPI; 1997-402555/37.
P-PSDB; AAW29212.
  Rhode PR, Jiao J,
  WO9728191-A1
  30-JAN-1997;
  31-JAN-1996;
   27-MAR-1998
  07-AUG-1997
  Synthetic
   645
                                 681
```

Key

Chavaillaz P, Jiao JJJ;

WPI; 2003-341126/32. P-PSDB; ABU72106. Novel major histocompatibility complex fusion complex having presenting peptide covalently linked to MHC molecule containing peptide-binding groove, used for suppressing immune response in multiple sclerosis,

Example 17; Fig 27; 126pp; English.

The invention relates to a major histocompatibility complex (MIC) fusion complex (I) comprising an MHC molecule that contains a peptide-binding groove, and a presenting peptide covalently (e.g. an antiganc peptide) linked to the MHC molecule, where (I) is capable of modilating the activity of a Teal. Also included are a DNA construct coining for the complex, where the MHC molecule is a class II MHC (e.g. mouse I-Ad or I-As, or human HLA-DRI (human leukocyte antigen-DRI)), a multivalent MHC fission complex comprising two or more linked complexes, identifying a peptide that can modilate the activity of T cells (lanvolving introducing into host cells cloning vectors that each contain the fusion complex and selecting host cells that express MGC fusion complex, and selecting host cells that express MGC fusion complex, and selecting host cells, a single recombinant expression vector comprising DNA that codes for the alpha and beta chains of the MHC fusion complex. Also included are inducing an immune expression of the MHC fusion complex. The DNA constructs can contain heterologous leader peptide sequences and Kozak sequence for efficient expression of the MHC fusion complex. The DNA constructs can contain heterologous leader peptide sequences and Kozak sequence for efficient casponse in a mammal (including vaccinating a fusion complex, or DNA sequence codprising a fusion complex, or DNA sequence codprising a fusion complex which is a single chain fusion molecule. The methods are useful for it en MFC protein, or DNA sequence codprising a peptide that is a T cell receptor (TCR) antagonist or partial agonist and is covalently linked to the MFC protein, or DNA sequence coding for the fusion complex which is a single chain fusion and eventual and several the tusion complex which is a single chain fusion and activity of T cells, inducing an immune response in a mammal agonist and several for its expression in a mammal agonism and mammal and mammal and mammal and mammal and mammal and mammal and mammal and mammal and mammal

Sequence 1385 BP; 316 A; 383 C; 399 G; 287 T; 0 U; 0 Other;

200 201 GGTGGAGGCTCCGAAAGGCATTTCGTGGTCCAGTTCAAGGGCGAGTGCTACTACACCAAC 260 81 AGCAGCCCCGGGACTGAGGGCGGAAACTCCATCTGCTTCTCGCCGTCGCTGGAGCACCCG 140 164 ------critaagtarcricricagicrigation 224 80 65 6 ATGGCTCTGCAGATCCCCCAGCCTCCTCTCAGCTGCTGGGTGGTGCTGATGGTGCTG 141 ATCGTGGTGTCCGGCAGCTGGACGAGGTGGGGGCTCACTAGTGCCCCGAGGCTCTGGA 165 GGGGGAAACTCCGAAAGGCATTTCGTGGTCCAGTTCAAGGGCGAGTGCTACACCAAC 21 ATGGCTCTGCAGATCCCCAGCCTCCTCTCAGCTGCTGTGGTGGTGCTGATGGTGCTG Score 607.8; DB 8; Length 1385; Pred. No. 1e-131; 0; Mismatches 57; Indels 21; 66 AGCAGCCCAAGGAC-----nery Match 63.9%; set Local Similarity 89.6%; stches 673; Conservative

320

GGGACGCAGCGCATACGGCTCGTGACCAGATACATCTACAACCGGGAGGAGTACGTGCGC

261

Single chain major histocompatibility complex comprising linked alpha and beta chains - useful for suppressing an immune response to an auto:Immune disease, e.g. multiple sclerosis, rheumatoid arthritis, diabetes mellitus, etc. 404 500 464 560 524 620 584 680 644 740 321 TACGACAGCGACGTGGGCGAGTACCGCGCGTGACCGAGCTGGGGCGGCCAGACGCCGAG 380 225 GGGACGCAGCGCATACGGCTCGTGACCAGATACATCTACAACCGGGAGGAGTACGTGCGC 284 645 GTCATGCTGGAGATGATCCCCTCATCAGGGAGAGGTCTACACCTGCCATGTGGAGGCCATCCC 285 TACGACAGCGACGTGGGGCGAGTACCGCGGGGGGACCGAGGCTGGGGCGGCCAGACGCCGAG 345 TACTGGAACAGCCAGCCGGAGATCCTGGAGCGAACGCGGGCCGAGGTGGACACGGCGTGC AGACACAACTACGAGGCCGGAGACCAGCACCTCCCTGCGGCGGCTTGAACAGCCCCAAT 525 GTGACAGATTTCTACCCAGCCAAGATCAAAGTGCGCTGGTTCAGGAATGGCCAGGAGGAG 381 TACTGGAACAGCCAGCCGGAGATCCTGGAGCGAAACGCGGGCCGAGGTGGACACGCGTGC 405 AGACACAACTACGAGGGCCGGAGACCAGCACCTCCCTGCGGGGGTTGAACAGCCCAAT GTCGCCATCTCCCTGTCCAGGACAGAGGCCCTCAACCACCACACACTCTGGTCTGTTCG 465 GTCGCCATCTCCCTGTCCAGGACAGAGCCCTCAACCACCACAACACTCTGGTCTGGTCG GTGACAGATTTCTACCCAGCCAAGATCAAAGTGCGCTGGTTCAGGAATGGCCAGGAGGAG ACAGTGGGGGTCTCATCCACACACGCTTATTAGGAATGGGGACTGGACCTTCCAGGTCCTG GTCATGCTGGAGATGACCCCTCATCAGGAAGAGGTCTACACCTGCCATGTGGAGCATCCC major histocompatibility complex; MHC; fusion complex; 705 AGCCTGAAGAGCCCCATCACTGTGGAGTGGA 735 741 AGCCTGAAGAGCCCCATCACTGTGGAGTGGA 771 Wong Location/Qualifiers 6. .1508 /\*tag= a Jiao J, Burkhardt M, AAT86988 standard; DNA; 1508 97WO-US001617 96US-00596387 Construction, major histoco SCT1 single chain gene; ss. (first entry) SCT1 single chain gene. (DADE-) DADE INT INC. WPI; 1997-402555/37. P-PSDB; AAW29213 27-MAR-1998 WO9728191-A1 30-JAN-1997; 31-JAN-1996; 07-AUG-1997. Rhode PR, Synthetic. 441 501 561 621 681 유 g ò 용 ઠે 셤 ઠે 8 ð 셤 ઠે 셤 ઠે 셤 ठ 셤

immune response;

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The invention relates to new single chain major histocompatibility complex (sc-MHC) class II complexes that comprise a peptide binding groove, and a modified class II beta 2 chain or covalantly linked imminoquobulin (Ig) light chain constant (CI) region. The MHC complexes are useful for detection and analysis of peptide ligands, pathogenic T-cells, for functional, cellular and molecular assays. They can be used to identify and isolate T cell receptor and/or MHC agonists and antagonists. They can also be used to calls involved in immune-related disorders. They can also be used to raise antibodies and to screen immune cells. It is also use in a method of suppressing an immune response in mammals. The sc-MHC complexes of suppressing an immune are in mammals. The sc-MHC complexes comprising modified class II beta 2 chains and/or Ig-C1 regions are soluble and provide enhanced yield. These MHC complexes also can contain single antigenic peptides readily isolated from expressing cells in significant quantities. The polyspecific MHC complexes also provide a means to detect cells expressing multiple target structures with a single complex. The present sequence represents a DNA encoding a single chain
   81 AGCAGCCCGGGACTGAGGGCGGAAACTCCATCTGCTTCTCGCCGTCGCTGGAGCACCCG 140
  201 GGTGGAGGCTCCGAAAGGCATTTCGTGGTCCAGTTCAAGGGCGAGTGCTACTACAAC 260
  261 GGGACGCAGCGCATACGGCTCGTGACATACATCTACAACCGGGAGGAGTACGTGCGC 320
  141 ATCGTGGTGTCCGGCAGCTGGGACGGGGGTGGGGGCTCACTAGTGCCCCCGAGGCTCTGGA
  105 gcrócrcaccicidaaárcaaccaaccicircirciraccacacaccccidaacccccc
   21 ATGGCTCTGCAGATCCCCAGCCTCCTCTCAGCTGCTGTGGTGGTGCTGATGGTGCTG
   6 aresercrecasarececaseerecerecreteasersersersersersarsers
  Single chain major histocompatibility complex class I complexes
                                   Major histocompatibility complex; MHC; single chain MHC; sc-MHC peptide binding groove; immunoglobulin; T cell receptor; immune immune-related disorder; antigenic peptide; fusion protein; ss
   Sequence 1508 BP; 337 A; 413 C; 441 G; 317 T; 0 U; 0 Other;
chain IAd/OVA 323-229 MHC fusion protein encoding
   Rhode PR, Acevedo J, Burkhardt M, Jiao J, Wong HC;
   Query Match
63.9%; Score 607.8; DB 2;
Best Local Similarity 89.6%; Pred. No. 1.1e-131;
Matches 673; Conservative 0; Mismatches 57;
  Example 1; Fig 1; 148pp; English.
  98WO-US021520.
   97US-00960190.
   (SUNO-) SUNOL MOLECULAR CORP
   WPI; 1999-418411/35.
  P-PSDB; AAY27111
  13-OCT-1998;
   29-OCT-1997;
  WO9921572-A1
  06-MAY-1999
  Synthetic.
    Single
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   200
  320
  380
  440
   200
   620
  644
  740
   680
   81 AGCAGCCCCGGGACTGAGGGCGGAAACTCCATCTGCTTCTCGCCGTCGCTGGAGCACCCGG 140
   344
   464
   524
   584
   164
   404
  704
   80
   65
   GGGACGCAGCGCATACGGCTCGTGACCAGATACATCTACAACGGGGAGGAGTACGTGCGC
  TACTGGAACAGCCAGCCGGAGATCCTGGAGCGGAACGCGGGCCGAGGTGGACACGGCGTGC
  AGACACAACTACGAGGGGCCGGAGACCAGCACCTCCCTGCGGCGCTTGAACAGCCCAAT
  GTGGCCATCTCCCTGTCCAGGACAGAGGCCCTCAACCACCACACACTCTGGTTCG
   GTGACAGATTTCTACCCAGCCAAGATCAAAGTGCGCTGGTTCAGGAATGGCCAGGAGGAG
   GTGACAGATITICTACCCAGCCAAGATCAAAGTGCGCTGGTTCAGGAATGGCCAGGAGGAG
   GTCATGCTGGAGATGACCCCTCATCAGGGAGAGGTCTACACCTGCCATGTGGAGCATCCC
  21 ATGGCTCTGCAGATCCCCAGCCTCCTCTCAGCTGCTGTGGTGGTGCTGATGGTGCTG
   141 ATCGTGGTGTCCCGCCAGCTGGGACGGGGGGCTCACTAGTGCCCCCGAGGCTCTGGA
   GGTGGAGGCTCCGAAAGGCATTTCGTGGTCCAGTTCAAGGGCGAGTGCTACTACAAC
  GGGACGCAGCGCATACGGCTCGTGACCAGATACATCTACAACCGGGAGGAGTACGTGCGC
  TACGACAGCGACGTGGGCGAGTACCGCGCGGTGACCGAGCTGGGGCGGCCCAGACGCCGAG
  TACTGGAACAGCCAGCCGGAGATCCTGGAGCGAACGCGGGCCCGAGGTGGACACGCGTGC
   AGACACAACTACGAGGGGCCGGAGACCAGCACCTCCCTGCGGCGGCGTTGAACAGCCCAAT
   ACAGTGGGGGTCTCATCCACACACCTTATTAGGAATGGGGACTGGACCTTCCAGGTCCTG
  Greangerigadarcacercareaggagagarcraeacercaecargagagarcee
  Gaps
  21;
  DB 2; Length 1508;
  Sequence 1508 BP; 337 A; 413 C; 441 G; 317 T; 0 U; 0 Other;
   The present sequence was used in the construction of major histocompatibility complex (MHC) fusion complexes
  ery Match 63.9%; Score 607.8; DB 2; Length : st Local Similarity 89.6%; Pred. No. 1.1e-131; tches 673; Conservative 0; Mismatches 57; Indels
  AGCCTGAAGAGCCCCATCACTGTGGAGTGGA 771
  Agccrigadadaccccarcacricregadrega 735
                       Example 17; Page 137-139; 217pp; English.
  B
  AAX89069 standard; DNA; 1508
  201
  261
  225
  321
   381
   345
   441
   405
   501
   465
   561
   525
  621
  681
  645
  741
  705
  AAX89069
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164

224

200

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21;

Indels

Length 1508;

284

321 TACGACAGCGACGTGGGCGAGTACCGCGGGGGACCGAGCTGGGGGCGGCCAGACGCGGAG 380

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(first entry)

14-SEP-1999

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500
  620
   680
  584
   644
  GTCATGCTGGAGATGACCCCTCATCAGGGAGAGGTCTACACCTGCCATGTGGAGCATCCC 740
  704
404
  464
  524
  Novel major histocompatibility complex fusion complex having presenting peptide covalently linked to MHC molecule containing peptide-binding groove, used for suppressing immune response in multiple sclerosis, allergies.
   345 TACTGGAACAGCCAGCCGGAGATCCTGGAGCGAACGCGGGCCGAGGTGGACACGGCGTGC
  465 Gredecarcrecererecasadadadadecereraaceacadaadacarcregregeres
  525 GTGACAGATTTCTACCCAGCCAAGATCAAAGTGCGCTGGTTCAGGAATGGCCAGGAGGAG
   ACAGTGGGGGTCTCCACCACACTTATTAGGAATGGGGACTGGACCTTCCAGGTCCTG
   585 ACAGTGGGGGTCTCATCCACACACAGCTTATTAGGAATGGGGACTGGACCTTCCAGGTCCTG
   645 GTCATGCTGGAGATGACCCCTCATCAGGGAGAGGTCTACACCTGCCATGTGGAGCATCCC
  381 TACTGGAACAGCCAGCCGGAGATCCTGGAGCGAACGCGGGCCGAGGTGGACACGGCGTGC
   441 AGACACAAACTACGAGGGCCGGAGACCAGCACCTCCCTGCGGCGGCGCTTGAACAGCCCAAŢ
  GTGACAGATTTCTACCCAGCCAAGATCAAAGTGCGCTGGTTCAGGAATGGCCCAGGAGGAG
  405 AGACACAACTACGAGGGCCGGAGACCAGCACCTCCCTGCGGCGGCTTGAACAGCCCAAT
  MHC; major histocompatibility complex; gene therapy; fusion complex; peptide-binding groove; T cell modulation; class II MHC; vaccine; autofinmune disorder; multiple sclerosis; rheumatoid arthritis; insulin-dependent diabetes mellitus; myasthenia gravis; immunogen; chronic allergy; mouse; ds; I-Ad; gene.
  Edwards AC;
   Mouse MHC I-Ad/Ova 323-339 synthetic gene SCT1.
  Grammer S,
  AGCCTGAAGAGCCCCATCACTGTGGAGTGGA 735
   AGCCTGAAGAGCCCCATCACTGTGGAGTGGA 771
   Rhode PR, Weidanz JA,
  ACA60743 standard; DNA; 1508 BP
  94US-00283302.
95US-00382454.
97US-00776084.
   06-JUL-2001; 2001US-00900379
   16-JUN-2003 (first entry)
  Chavaillaz P, Jiao JJJ;
  (DADE-) DADE INT INC.
  WPI; 2003-341126/32.
   P-PSDB, ABU72107.
   US2002198144-A1
  29-JUL-1994;
01-FEB-1995;
17-JAN-1997;
  26-DEC-2002
  Synthetic.
  741
   705
  501
   621
  681
   ACA60743,
  Wong HC,
   561
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The invention relates to a major histocompatibility complex (WHC) fusion complex (I) comprising an MHC molecule that contains a peptide-binding grower, and a presenting peptide covalently (e.g. an antiganic peptide) inked to the MHC molecule, where (I) is capable of modulating the tenderouse, where the MHC molecule is a class II MHC (e.g. mouse I-Ad or I complex, where the MHC molecule is a class II MHC (e.g. mouse I-Ad or I complex, where the MHC molecule is a class II MHC (e.g. mouse I-Ad or I complex, where the MHC molecule is a class II MHC (e.g. mouse I-Ad or I complex complex complex to modulate the activity of T cells (involving introducing peptide that can modulate the activity of T cells (involving introducing culturing the host cells under conditions suitable for expression of the complex who selecting host cells in a single recombinant complex that modulate the activity of T cells), a single recombinant complex that modulate the activity of T cells), a single recombinant complex that codes for a T cell costimulatory factor and the alpha and beta chains of the MHC fusion complex. The DNA constructs can contain cherrologous leader peptide sequences and kozak sequence for efficient expression of the fusion complex. Also included are inducing an immune complex, by administering DNA sequence comprishing a nemmal against a targeted clisorder, by administering by sequence comprishing an expression vector, encoding for a fusion complex which is a single chain fusion complex, or the mammal a DNA sequence comprishing an expression vector, encoding to the fusion complex which is a single chain fusion complex or the mammal and suppressing an immune response in a mammal including vaccinating a peptide that is a T cell receptor (TCR) antegonist or partial configuration and suppressing an immune response in a mammal. The disorders and or suppressing an immune response in a mammal. The disorders included an autolimmune disorder configured in a mammal including arranting a single chain a subsetution or humannes an au
  200
  320
   81 AGCAGCCCCGGGACTGAGGGCGGAAACTCCATCTGCTTCTCGCCGTCGCTGGAGCACCCG 140
   224
   284
  321 TACGACAGCGACGTGGGCGAGTACCGCGCGTGACCGAGCTGGGGCGGCCAGACGCCGAG 380
   345 TACTIGGAACAGCCAGCCGGAGATCCTGGAGCGAACGCGGGCCGAGGTGGACACGGCGTGC 404
   6 Aresercriscasariceceasecricererereaseriscriscasississississis es
   GGGACGCACGCATACGGCTCGTGACCAGATACATCTACAACCGGGAGGAGTACGTGCGC
  141 ATCGTGGTGTCCGGCAGCTGGACGGAGGTGGGGCTCACTAGTGCCCCGAGGCTCTGGA
   261 GGGACGCAGCGCATACGGCTCGTGACCAGATACATCTACAACCGGGAGGAGTACGTGCG
   381 TACTGGAACAGCCGGAGATCCTGGAGCGAACGCGGGCCGAGGTGGACACGCGTGC
  21 ATGCCTCTGCAGATCCCCAGCCTCCTCTCAGCTGCTGTGGTGGTGGTGGTGGTGCTG
  21,
   Query Match 63.9%; Score 607.8; DB 8; Length 1508; Best Local Similarity 89.6%; Pred. No. 1.1e-131; Matches 673; Conservative 0; Mismatches 57; Indels 21;
   Sequence 1508 BP; 337 A; 413 C; 441 G; 317 T; 0 U; 0 Other;
                   Example 17; Fig 28; 126pp; English
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Wong HC, Rhode PR, We
Chavaillaz P, Jiao J;
   (DADE-) DADE INT INC.
  WPI; 1996-129343/13.
P-PSDB; AAR98907.
  31-JUL-1995;
  29-JUL-1994;
01-FEB-1995;
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   WO9604314-A1
   15-FEB-1996.
  81
  261
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   요
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   ò
   620
  680
   584
  644
  524
  6. 86
/*tag= b
//label= I-Ad beta chain leader
/note= "murine MHC class II I-Ad gene beta chain leader
   MHC; major histocompatibility complex; PCR; polymerase chain reaction; T cell activity modulator; antagonist; immune disorder; allergy; multiple sclerosis; insulin-dependent diabetes mellitus; rheumatoid arthritis; myasthenia gravis; ds.
   GTGACAGATTTCTACCCAGCCAAGATCAAAGTGCGCTGGTTCAGGAATGGCCAGGAG
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/note= "murine MHC class II I-Ad gene alpha-2 domain"
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   GTCGCCATCTCCCTGTCCAGGACAGAGCCCTCAACCACCACACACTCTGGTCTGTTCG
   GTGACAGATTTCTACCCAGCCAAGATCAAAGTGCGCTGGTTCAGGAATGGCCAGGAGGAG
  ACAGTGGGGGTCTCATCCACACACACTTATTAGGAATGGGGACTGGACCTTCCAGGTCCTG
  GTCATGCTGGAGATGACCCCTCATCAGGGAGGGTCTACACCTGCCATGTGGAGCATCCC
   /label= I-Ad_beta2
/note= "murine MHC class II I-Ad gene beta-2 domain"
735. .806
   /*tag= e
/label= I-Ad_betal
hotte= "murine MHC class II I-Ad gene beta-1 domain"
453. .71
  Vector SCE1-derived single chain gene encoding MHC fusion complex.
  /*tag= c
/labbl= OVA 323-339
/note= "chicken ovalbumin residues 323-339"
138. .167
   /*tag= g
/note= "24 residue peptide linker"
807. .1067
   /*tag= d
/note= "10 residue linker peptide"
  771
   705 AGCCTGAAGACCCCCATCACTGTGGAGTGGA 735
   741 AGCCTGAAGAGCCCCATCACTGTGGAGTGGA
   Location/Qualifiers
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AAT17588 encodes a murine MHC fusion complex capable of modulating T cell activity encoded by the vector SCEI. The MHC fusion complex comprises at least one MHC molecule containing a peptide-binding groove and a presenting peptide covalently linked to the MHC molecule and opt. a transmembrane domain. DNA encoding a MHC fusion complex may be cloned transmembrane domain. DNA encoding a MHC fusion complex may be cloned controlled to express that modulate, pref. antagonise, T cell activity. DNA encoding a MHC fusion complex or a single chain fusion concleve may be used to vaccinate a mammal against a targeted disorder. The fusion complexes may be used to suppress an immune response in an animal suffering from an immune disorder e.g. multiple sclerosis, insulin chapter and pets such a seats and dogs. The MHC fusion complexes may also be used in the treatment of chronic allergies. The complexes may also be used in the treatment of chronic allergies. The complexes may also be used in the treatment of the produced such that they contain a single antigenic peptide including one of known structure, additionally a wide range of peptides can be contain a single antigenic peptide including one contain a single antigenic peptide can be contain a single antigenic peptides can be contain a single antigenic peptide including one contain a single antigenic peptide can be
  200
  164
   260
  104
   224
  65
  6 aresercrecasarceceasecrecrecrereasersersersersersers
  141 ATCGTGGTGTCCGGCAGCTGGACGGAGGTGGGGGCTCACTAGTGCCCCGAGGCTCTGGA
  105 gcrócrcacecréanárcaacéanécréércérécracceanégéséséséanéceses
   201 GGTGGAGGCTCCGAAAGGCATTTCGTGGTCCAGTTCAAGGGCGAGTGCTACTACACCAAC
   GGGACGCAGCGCATACGGCTCGTGACCAGATACATCTACAACCGGGAGGAGTACGTGCGC
  66 agcacccaaggac-----critaagrarcricacgcrgracac
  21 Areacreracharecechacerecrererenacracraseracranaracras
  Major histocompatability complex fusion complex for modulating T cell activity - used in the treatment of immune disorders, e.g. multiple sclerosis, IDDM and rheumatoid arthritis.
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/labba I-Ad_alpha2
/note= "murine MHC class II I-Ad gene alpha-2 domain"
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63.7%; Score 606.2; DB 2; Length 1382;
Best Local Similarity 89.5%; Pred. No. 2.4e-131;
Matches 672; Conservative 0; Mismatches 58; Indels 21;
   Sequence 1382 BP; 320 A; 374 C; 404 G; 284 T; 0 U; 0 Other;
  Edwards AC;
  Grammer S,
  Example 17; Fig 29; 210pp; English.
  Weidanz JA,
   /*tag= j
/note= "EE tag"
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   94US-00283302.
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   94US-00283302.
  Wong HC, Rhode PR, We
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   (DADE-) DADE INT INC.
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   31-JUL-1995;
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   WO9604314-A1
   01-PEB-1995;
  misc_feature
   15-PEB-1996.
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  561 GTGACAGATTTCTACCCAGCCAAGATCAAAGTGCGCTGGTTCAGGAATGGCCAGGAGGAG 620
  345 TACTGGAACAGCCAGCCGGAGATCCTGGAGCGAACGCGGGCCGAGGTGGACACGGCGTGC 404
  405 AGACACAACTACGAGGGCCGGAGACCAGCACCTCCCTGCGGGGGGGTTTGAACAGCCCAATT 464
   GTCGCCATCTCCCTGTCCAGGACAGAGGCCCTCAACCACACAACACTCTGGTCTGTTCG 560
  524
   680
   740
225 GGGACGCAGCGCATACGGCTCGTGACCAGATACATCTACAACCGGGAGGAGTACGTGCGC 284
                           321 TACGACAGCGACGTGGGCGAGTACCGCGGGGGACCGAGCTGGGGCGGCCAGACGCCGAG 380
  285 TACGACAGCGACGTGGCCGAGTACCGCGGTGACCGAGCTCGGGCGGCCAGACGCCGAG 344
   381 TACTGGAACAGCCAGCCGGAGATCCTGGAGCGAACGCGGGCCGAGGTGGACACGGCGTGC 440
  525 GIGACAGAITICIACCCAGCCAAGAICAAAGIGCGCIGGIICAGGAAIGGCCAGGAGGAG 584
  585 ACAGIGGGGICICATCCACACACTIATIAGGAAIGGGGACIGGACTICCAGGICCIG 644
   645 Greargeregagargaceerearcagagagagaereracaceraceargragageareee 704
   /label- I-Ad beta chain leader
/note= "murine MHC class II I-Ad gene beta chain leader
sequence"
   MHC; major histocompatibility complex; PCR; polymerase chain reaction; T cell activity modulator; antagonist; immune disorder; allergy; multiple sclerosis; insulin-dependent diabetes mellitus; rheumatoid arthritis; myasthenia gravis; ds.
  441 AGACACAACTACGAGGGCCGGAGACCAGCACCTCCCTGCGGGGGCGCTTGAACAGCCCAAT
   GTCATGCTGGAGATGACCCCTCATCAGGGAGAGGTCTACACCTGCCATGTGGAGCATCCC
   ACAGTGGGGGTCTCCATCCACACGCTTATTAGGAATGGGGACTGGACCTTCCAGGTCCTG
  /*tag= e
/label= I-Ad_beta1
/note= "murine MHC class II I-Ad gene beta-1 domain"
/*tag= f
  Vector SSC1-derived single chain gene encoding MHC fusion complex
   'trag= c
label= OVA 323-339
'note= "chicken ovalbumin residues 323-339"
   138. .167
/*tag= d
/otts= 110 residue linker peptide"
168. .452
  705 AGCCTGAAGAGCCCCATCACTGTGGAGTGGA 735
   741 AGCCTGAAGAGCCCCATCACTGTGGAGTGGA 771
  Location/Qualifiers
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  (first entry)
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  .137
   *tag=
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  26-SEP-1996
  misc_feature
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  Synthetic
   501
  681
   AAT17586;
   621
  Key
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AAT17586 encodes a murine MHC fusion complex capable of modulating T cell activity encoded by the vector SSC1. The MHC fusion complex comprises at least one MHC molecule containing a peptide-binding groove and a presenting peptide covalently linked to the MHC molecule and opt. a transmembrane domain. DNA encoding a MHC fusion complex may be cloned into a host cell to express the complex. The transformed cells may then cet into a host cell to express that modulate, pref. antegonise, T cell activity. DNA encoding a MHC fusion complex or a single chain fusion conjecule may be used to vaccinate a mammal against a targeted disorder. The fusion complexes may be used to suppress an immune response in an animal suffering from an immune disorder e.g. multiple sclerosis, insulin chapendent diabetes mellitus, rheumacioid arthritis, myasthenia gravis or chronic allergies. The complexes may also be used in the treatment of chronic allergies. The complexes may also be used in the treatment of chronic allergies. The complexes may also be used in the treatment of chronic allergies that they contain a single antigenic peptide including one of known structure, additionally a wide range of peptides can be complexed to the contain a single antigenic peptide including one complexes to the contain a single antigenic peptide including one complexes to the contain a single antigenic peptide can be contained for T cell interaction
  140
  141 ATCGTGGTGTCCCGGCAGCTGGAACGGAGGTGGGGCTCACTAGTGCCCCGAGGCTCTGGA 200
   65
  6 AIGGCTCTGCAGATCCCCAGCTCCTCTCAGCTGCTGCTGGTGGTGCTGATGGTGCTG
  21 ATGGCTCTGCAGATCCCCCAGCTCCTCTCAGCTGCTGTGGTGGTGCTGATGGTGCTG
  /*tag= h
/label= I-Ad_alphal
/note= "murine MHC class II I-Ad gene alpha-2 domain"
1068. .1352
  Major histocompatability complex fusion complex for modulating T cell activity - used in the treatment of immune disorders, e.g. multiple sclerosis, IDDM and rheumatoid arthritis.
   gene alpha-2 domain"
   Gaps
/label= I-Ad_beta2
/note= "murine MHC class II I-Ad gene beta-2 domain"
/35. .806
   21;
   Query Match 63.7%; Score 606.2; DB 2; Length 1385; Best Local Similarity 89.5%; Pred. No. 2.4e-131; Matches 672; Conservative 0; Mismatches 58; Indels 21;
   Sequence 1385 BP; 316 A; 384 C; 398 G; 287 T; 0 U; 0 Other;
  Grammer S, Edwards AC;
   /*tag= g
/note= "24 residue peptide linker"
307. .1067
   class II I-Ad
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/label= I-Ad_alpha2
/note= "murine MHC cl
1353. .1382
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404
   GTGACAGATTTCTACCCAGCCAAGATCAAAGTGCGCTGGTTCAGGAATGGCCAGGAGGAG 620
   704
   GGGACGCAGCCATACGCCTCGTGACCAGATACATCTACAACCGGGAGGAGTACGTGCGC 284
   6..86
/*tag= b
/labal= I-Ad beta chain leader
/note= "murine MHC class II I-Ad gene beta chain leader
  MHC; major histocompatibility complex; PCR; polymerase chain reaction; T cell activity modulator; antagonist; immune disorder; allergy; multiple sclerosis; insulin-dependent diabetes mellitus; rheumatoid arthritis; myasthenia gravis; ds.
  345 TACTGGAACAGCCAGCCGGAGATCCTGGAGCGAACGCGGGCCGAGGTGGACACGCGGCGTGC
  405 AGACACAACTACGAGGGCCCGGAGACCAGCACCTCCCTGCGGGGGGCTTGAACAGCCCAAT
  525 GTGACAGATTTCTACCCAGCCAAGATCAAAGTGCGCTGGTTCAGAATGGCCAGGAGGAG
   645 ercarecresagareaccercarcaseagagaercracacereceareresagearece
                                   GGTGGAGGCTCCGAAAGGCATTTCGTGGTCCAGTTCAAGGGCGAGTGCTACTACACCAAC
   GGGACGCAGCGCATACGGCTCGTGACCAGATACATCTACAACCGGGAGGAGTACGTGCGC
  TACGACAGCGACGTGGGCGAGTACCGCGCGGGGAGCTGGGGCGGCCGAGACGCCGAG
   TACTGGAACAGCCAGCCGGAGATCCTGGAGCGAACGCGGGCCGAGGTGGACACGCCGTGC
   AGACACAACTACGAGGGGCCGGAGACCAGCACCTCCCTGCGGCGGCGCTTGAACAGCCCAAT
   GTCGCCATCTCCCTGTCCAGGACAGAGGCCCTCAACCACCACACACTCTGGTCTGTTCG
   621 ACAGTGGGGGTCTCATCCACAGCTTATTAGGAATGGGGACTGGACCTTCCAGGTCCTG
  GTCATGCTGGAGATGACCCCTCATCAGGAGGAGGTCTACACCTGCCATGTGGAGCATCCC
   Vector SCT1-derived single chain gene encoding MHC fusion complex
  /*tag= c
/label= OVA 323-339
/note= "chicken ovalbumin residues 323-339"
   741 AGCCTGAAGAGCCCCATCACTGTGGAGTGGA 771
  705 AGCCTGAAGAGCCCCATCACTGTGGAGTGGA 735
   Location/Qualifiers
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   Synthetic
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   681
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  261
  225
  321
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   501
   561
  AAT17587;
   Key
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AAT17587 encodes a murine MHC fusion complex capable of modulating T cell activity encoded by the vector SCT1. The MHC fusion complex comprises at least one MHC molecule containing a peptide-binding groove and a presenting peptide covalently linked to the MHC molecule and opt. a creamembrane domain. DNA encoding a MHC fusion complex may be cloned transmembrane domain. DNA encoding a MHC fusion complex may be cloned creativity. DNA encoding a MHC fusion complex may be used to identify peptides that modulate, pref. antagonise, T cell activity. DNA encoding a MHC fusion complex or a single chain fusion complexes may be used to suppress an immune response in an canimal suffering from an immune disorder e.g. multiple sclerosis, insuling chronic allergies. The complexes may also be used in the treatment of chronic allergies. The complexes may also be used in the treatment of chronic allergies. The complexes may also be used in the treatment of chronic allergies. The complexes may also be used in the treatment of chronic allergies. The complexes may also be used in the treatment of chronic allergies. The complexes may also be used in the treatment of chronic allergies. The complexes may also be used in the treatment of chronic allergies. The complexes may also be used in the treatment of chronic allergies and dogs. The MHC fusion complexes can be produced such that they contain a single antigenic peptide including one of produced for T cell interaction
   /*tag= j
/label= I-Ad_alpha-TM
/note= "murine MHC class II I-Ad gene alpha-transmembrane
domain"
  Major histocompatability complex fusion complex for modulating T cell activity - used in the treatment of immune disorders, e.g. multiple sclerosis, IDDM and rheumatoid arthritis.
   /*tag= h
/label= I-Ad alpha1
/note= "murine MHC class II I-Ad gene alpha-2 domain"
1068. .1352
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/label= I-Ad_alpha2
/label= "murine MHC class II I-Ad gene alpha-2 domain"
|353. .1505
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/label= I-Ad_betal
/note= "muxine MHC class II I-Ad gene beta-1 domain"
453. .734
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   beta-2
  Sequence 1508 BP; 337 A; 414 C; 440 G; 317 T; 0 U; 0 Other;
  Wong HC, Rhode PR, Weidanz JA, Grammer S, Edwards AC;
Chavaillaz P, Jiao J;
  /*tag= f
/label= I-Ad beta2
/note= "muxine MHC class II I-Ad gene }
735. .806
38. .167
*tag= d
note= "10 residue linker peptide"
  *tag= g
/note= "24 residue peptide linker"
807. 1067
  Example 17; Fig 28; 210pp; English.
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89.5%;
   95WO-US009816
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95US-00382454
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  168.
   (DADE-) DADE INT INC.
   WPI; 1996-129343/13.
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Best Local Similarity
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  WO9604314-A1
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   440
  500
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  284
  584
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   105 gcrecrcacecreaaarcaaceaacerecrecrecraeceaaeseseseseseseseseses
   345 TACTGGAACAGCCAGCCGGAGATCCTGGAGCGAACGCGGGCCGAGGTGGACACGGCGTGC 404
  6 ATGGCTCTGCAGATCCCCAGCCTCCTCTCAGCTGCTGTGGTGGTGCTGATGGTGCTG 65
  165 GGGGGAAACTCCGAAAGGCATTTCGTGGTCCAGTTCAAGGGCGAGTGCTACTACACCAAC
   225 GGGACGCAGCGCATACGGCTCGTGACCAGATACATCTACAACCGGGAGGAGTACGTGGCGC
   321 TACGACACGCGACGTGGGCGAGTACCGCGCGGTGACCGAGCTGGGGCGGCCAGACGCCGAG
  381 TACTGGAACAGCCAGCCGGAGATCCTGGAGCGAACGCGGGCCGAGGTGGACACGGCGTGC
  441 AGACACAACTACGAGGGCCGGAGACCAGCACCTCCCTGCGGCGGCTTGAACAGCCCAAT
   681 GTCATGCTGGAGATGACCCCTCATCAGGGAGAGGTCTACACCTGCCCATGTGGAGCATCCC
58; Indels 21; Gaps
   741 AGCCTGAAGAGCCCCATCACTGTGGAGTGGA 771
  705 AGCCTGAAGAGCCCCATCACTGTGGAGTGGA 735
0; Mismatches
itches 672, Conservative
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**BLOSUM62** 

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Aavi2068 Murine IA

Aac104269 Wector SC

Aat1588 Vector SC

Aat86989 SCEI sing

Aca60744 Wouse MHC

Aat17586 Wector SS

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Aca60742 Wouse MHC

Aat17597 Vector SC

Aat8698 SCTI sing

Aca60742 Mouse MHC

Ad175986 Marker ge

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Acces
   Recombinant protein; alpha chain; beta chain; MHC; immunoglobulin; major hierocompatibility complex; FC region; antigen; T lymphocyte; immunostimulant; vaccine; infection; tumour; ss.
  DNA encoding a fusion protein comprising a beta chain of MHC.
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1151
  AAF55099;
  Aaf55099 DNA encod
Aat04262 Hybrid IA
Adg31225 I-Ab(beta
   May 31, 2006, 23:04:55 ; Search time 714 Seconds (without alignments) 4482.160 Million cell updates/sec
  Pred. No. is the number of results predicted by chance to have a score greater than or equal to the score of the result being printed, and is derived by analysis of the total score distribution.
  US-10-048-116B-6
1620
1 MALQIPSLLLSAAVVVLMVL......LKWKLQALKKKLAQHHHHHH 306
   Description
   10489840
                                 GenCore version 5.1.8
Copyright (c) 1993 - 2006 Biocceleration Ltd.
   nucleic search, using frame_plus_p2n model
   hits satisfying chosen parameters:
   5244920 segs, 3486124231 residues
  SUMMARIES
   -processing: Minimum Match 0%
Maximum Match 100%
Listing first 45 summaries
  AAF55099
AAT04262
ADQ31225
   Xgapop 10.0 , Xgapext
Ygapop 10.0 , Ygapext
Fgapop 6.0 , Fgapext
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genesequ200s: *
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   geneseqn2004bs:*geneseqn2005s:*
   geneseqn2006s:
  ü
  Geneseq_8:*
geneseqn1980s:*
  .mum DB seq length: 0
.mum DB seq length: 200000000
  17 2 2
  8
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Match Length
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base:

100.0 77.5 76.2

1620 1255.5 1235

Score

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(CNRS ) CNRS CENT NAT RECH SCI.
   Glaichenhaus N, Malherbe L;
  WPI; 2001-182944/18.
P-PSDB; AAB67481.
```

New soluble recombinant protein, useful e.g. as immunostimulant, comprises dimeric major histocompatibility complex molecule fused to immunoglobulin Fc region.

Example 1; Page 34-35; 43pp; French

The specification describes soluble recombinant proteins that comprise at least a dimer formed from the alpha and beta-chains of MHC (major histocompatibility complex) Class I and II molecules in which at least one chain has, attached to its C-terminus, at least part of the Fc region of an immunoglobulin. The recombinant proteins, when linked to an antigenied, part used to count and/or purify antigen-reactive T lymphocytes and to characterize their phenotype, e.g. in preclinical evaluation of vaccines. They are also used as immunostimulants, particularly for vaccine development (against infections and tumours), to count and determine phenotype of autoreactive T cells in subjects with, or at risk of developing, autoimmune diseases, e.g. for staging or evaluating treatments, and (to purify and/or enrich Ag-reactive T cells from cell cultures or patient samples, for use in subsequent curative or preventative cellular therapy. The present sequence encodes a recombinant protein of the invention, comprising a beta chain of MHC molecules

Sequence 921 BP; 214 A; 265 C; 286 G; 156 T; 0 U; 0 Other;

|               | 921       | 306      | 0               | 0                 | 0        | 0     |
|---------------|-----------|----------|-----------------|-------------------|----------|-------|
|               | Length:   | Matches: | Conservative:   | Mismatches:       | Indels:  | Gaps: |
|               | 6.68e-144 | 1620.00  | 100.00          | 100.04            | 100.04   | S     |
| nment Scores: | . No.:    | :0       | ent Similarity: | Local Similarity: | y Match: |       |

|                                                                        |              |                                                                                |                                                                  | 0                                     | 0                                                                |                                                                         | 0 0                                                              | 0 0                                                                                              | 0 0                                                                             |
|------------------------------------------------------------------------|--------------|--------------------------------------------------------------------------------|------------------------------------------------------------------|---------------------------------------|------------------------------------------------------------------|-------------------------------------------------------------------------|------------------------------------------------------------------|--------------------------------------------------------------------------------------------------|---------------------------------------------------------------------------------|
|                                                                        |              | 20                                                                             |                                                                  | 60                                    | 80                                                               | 300                                                                     | 120<br>360                                                       | 140                                                                                              | 160                                                                             |
|                                                                        |              | ]_Fe                                                                           | Pro H                                                            | 66—61<br>66—24                        | rThrasn<br>       <br> CACCAAC                                   | Arg<br> - -                                                             | 63<br>                                                           | 7<br>1<br>1<br>1<br>1<br>1<br>1<br>1<br>1<br>1<br>1<br>1<br>1<br>1<br>1<br>1<br>1<br>1<br>1<br>1 | ArgHishsnTyrGluGlyProGluThrSerThrSerLeuArgArgLeuGluGlnProAsn<br>                |
|                                                                        |              | Va   -                                                                         | H158                                                             | Ser<br>TCT                            | H-F                                                              | Val                                                                     | 4 – 8<br>- 8<br>- 8<br>- 8                                       | 4 – 8<br>8 – 8<br>8 – 8                                                                          | 91—17                                                                           |
|                                                                        |              | MetalaLeuGlnIleProSerLeuLeuLeuSeralaalaValValValUeuMetValLeu<br>amponementalan | 1610<br>                                                         |                                       | E S                                                              | GlythrGlnargileargLeuValthrargtyriletyrasnarggluGlutyrValarg<br>        | TyraspseraspvalglygluTyrargalavalThrGluLeuGlyargFroAspAlaglu<br> | TyrTrpAgnSerGlnProGluIleLeuGluArgThrArgAlaGluValAspThrAlaCys<br>                                 |                                                                                 |
|                                                                        |              | <u>1</u> _[                                                                    |                                                                  |                                       | G T                                                              | 64—69<br>64—69                                                          | 30-<br>30-<br>30-                                                | JABI<br>GACA                                                                                     | [6]<br>[7]                                                                      |
|                                                                        |              | 1.va                                                                           | ose<br>                                                          | 7-15<br>83-15<br>84-15                | G                                                                | - 3g<br>- 3g<br>- 3g<br>- 3g                                            | X-A-S                                                            | uVa<br>                                                                                          | g-green                                                                         |
| 921<br>0<br>0<br>0                                                     |              | 1Va                                                                            |                                                                  | ay — A                                | <u>6</u>                                                         | AH-CC-CC-CC-CC-CC-CC-CC-CC-CC-CC-CC-CC-CC                               | 15—15<br>60—15<br>60—15                                          | 15 — 86<br>16 — 86<br>16 — 86                                                                    | 9Ar<br>                                                                         |
|                                                                        |              | a<br>                                                                          | lese<br>                                                         |                                       | 12 <u>—</u> 25                                                   | 8=3<br>5=4                                                              | d Lee                                                            | 48—98<br>14—098                                                                                  | # - S                                                                           |
| : :                                                                    |              | [AB]                                                                           |                                                                  |                                       | <u>3</u>                                                         |                                                                         | 6 = 5<br>6 = 5                                                   | 3==¥                                                                                             | 7. L                                                                            |
| s s s s s s s s s s s s s s s s s s s                                  |              | erA]                                                                           |                                                                  | <u> </u>                              | Lup<br>G                                                         |                                                                         | GA T                                                             | EGT<br>SAAC                                                                                      | irse<br>T                                                                       |
| Length:<br>Matches:<br>Conservative<br>Mismatches:<br>Indels:<br>Gaps: | _            | ens<br>                                                                        |                                                                  | 9-15<br>10-15<br>10-15                | a1G<br>TCC                                                       | rg H                                                                    | 1aV<br>                                                          | AGC<br>HGC<br>HGC<br>HGC<br>HGC<br>HGC<br>HGC<br>HGC<br>HGC<br>HGC<br>H                          | erT]                                                                            |
| Lengt)<br>Match<br>Conse<br>Mismal<br>Indel                            | (1-921)      | eul<br>7—7                                                                     | ans — E                                                          | 8—5g                                  | a1   1                                                           | Š=P                                                                     | 49—98<br>                                                        | 9 — 6<br>1<br>1<br>1<br>1<br>1<br>1<br>1                                                         | hrs<br>                                                                         |
|                                                                        | ÷            |                                                                                | 4   1   1   1   1   1   1   1   1   1                            | 90-80<br>                             | hev<br>TCG                                                       | A I I                                                                   | AT'S                                                             | 13—F                                                                                             | Tale                                                                            |
|                                                                        | 660          | eric<br>Series                                                                 |                                                                  | 6-59<br>6-59                          | 11.8P                                                            | 14—e                                                                    | HuT<br>HAGT                                                      | Hall                                                                                             | 0-5<br>2-5<br>2-5                                                               |
| .68e-144<br>620.00<br>00.0%<br>00.0%                                   | AAF55099     | Silling<br>Silling                                                             |                                                                  |                                       | F                                                                | 19—55<br>—55                                                            | 4 – 8<br>– 8<br>– 8                                              | 6-8<br>8-8                                                                                       | 1.<br>1.<br>1.<br>1.<br>1.<br>1.<br>1.<br>1.<br>1.<br>1.<br>1.<br>1.<br>1.<br>1 |
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| 5 1000                                                                 | ×            |                                                                                |                                                                  | 5<br>5<br>5<br>5<br>5<br>5            | 8<br>                                                            | 7<br>- 1<br>- 1<br>- 1<br>- 1<br>- 1<br>- 1<br>- 1<br>- 1<br>- 1<br>- 1 | Pap<br>SAC                                                       | 3e r<br>6CC                                                                                      | <u> </u>                                                                        |
| ä                                                                      | (1-306)      | j—č                                                                            |                                                                  | 31—83<br>31—83<br>31—83               | 3<br>3<br>3<br>5<br>5<br>5                                       | 3<br>3<br>3<br>3<br>3<br>3<br>3<br>3                                    | Ser/                                                             | A-B                                                                                              | 7 - S                                                                           |
| <u>r</u>                                                               | (1-:         | 4 - K                                                                          | Serí                                                             | 8 = 5<br>1 = 9                        | 8<br>8<br>8<br>8                                                 | 47—5<br>7—5                                                             | ABD<br>SAC                                                       | <u>g</u> g                                                                                       | H = 5                                                                           |
| nt Scores: ).: Similarity: al Similarity itch:                         |              | A Section                                                                      | SerSerProdiyThrdluGlyGlyAenSerlleCyePheSerProSerLeuGluHisPro<br> |                                       | GlyGlyGlyGerGluArgHisPhevalValGlnPheLysGlyGluCysTyrTyrThrAsn<br> | GlythrGlnargileargLeuvalthrargtyriletyrasnargGluGlutyrValarg<br>        | 11-14-14-14-14-14-14-14-14-14-14-14-14-1                         | 11-14<br>12-14                                                                                   | Arg<br>                                                                         |
| Scores<br>milari<br>Simil<br>h:                                        | 16B          | -i -                                                                           |                                                                  |                                       | 61                                                               | 81                                                                      | 101                                                              | 121                                                                                              | 141                                                                             |
| No.:<br>No.:<br>nt Sim                                                 | 8-1          |                                                                                |                                                                  | -                                     | 7                                                                | 7                                                                       | 3 1                                                              | 3 1                                                                                              | н ч                                                                             |
| 5 5 5 5 5 5 5 5 5 5 5 5 5 5 5 5 5 5 5                                  | J-048-116B-6 |                                                                                |                                                                  |                                       |                                                                  |                                                                         |                                                                  |                                                                                                  |                                                                                 |
| ⊼ ∙ ਹ ਹੈ ਂ ≻                                                           |              |                                                                                |                                                                  |                                       |                                                                  |                                                                         |                                                                  |                                                                                                  |                                                                                 |

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   220
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  240
   GICATGCTGGAGATGACCCCTCATCAGGGAGAGGTCTACACCTGCCATGTGGAGCATCCC 720
   260
   280
   LystystyshanaladlnieutysTrptysteuGlnalaLeutystysteualadln 300
   ThrvalGlyvalSerSerThrGlnLeuIleArgAsnGlyAspTrpThrPheGlnValLeu
   221 ValMetLeuGluMetThrProHisGlnGlyGluValTyrThrCysHisValGluHisPro
   SerLeuLysSerProlleThrValGluTrpArgAlaGlnSerGluSerAlaArgSerLys
   /*tag= e
/note= "binding site for primer #331 (see AAT04261)"
complement(158. .212)
/*tag= f
/note= "primer #332 binding site"
complement(199. .250)
   /*tag= b
/note= "binding site for primer #261 (see AAT04260)"

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    complement (45. . 74)

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371.389
  Polymerase chain reaction; PCR; primer; amplify; major histocompatability complex; MRC; T-cell receptor; TCR; autoimmune disease; immunodeficiency disease; immune response; immunoproliferation disease; graft-host rejection; therapy; ss.
   /*tag= c
/product= "hybrid IA beta chain"
61. .141
   /*tag= d
/note= "leader region"
complement(119. 172)
  Location/Qualifiers
  нівнівнівнівнів 306
   BP
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   16-APR-1996 (first entry)
  Hybrid IA beta chain gene.
   61. .828
   misc feature
   primer_bind
  primer_bind
  primer_bind
   sig_peptide
  primer_bind
   primer_bind
  301 ]
  AAT04262;
   Synthetic
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  241
   601
   261
  281
                  481
   199
  901
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   This sequence represents a hybrid IA beta chain gene, containing the chicken ovalbumin peptide (COVA). This sequence was used in the construction of a hybrid IA alpha beta dimer. The encoded protein (plAd-OVA) was found to be more stable than the IA alpha beta dimer. The stability was decreased by the addition of a MHC groove specific binding peptide (e.g. see AAR82527, AAR82528 and AAR82531), compared to an increase seen on the addition of a MHC binding peptide to IB k/d-MCC. These complexes may be used to regulate an immune response. The complexes are capable of being recognised by a TCR alone or in combination with additional MHC proteins. These complexes are useful for therapeutic purposes and experimental purposes. They can also be used as reagents the treatment of diseases including autoimmune diseases, immunodeficiency diseases, immunoproliferation diseases, and graft-host rejection
  61 ATGGCTCTGCAGATCCCCAGCCTCCTCTCAGCTGCTGTGGTGGTGCTGATGGTGCTG 120
  Peptide-MHC complex comprising antigenic peptide, linker and MHC segment - useful as reagents for the treatment of diseases including auto-immune diseases, immuno-stimulatory diseases or graft-host rejection.
   MetAlaLeuGlnIleProSerLeuLeuLeuSerAlaAlaValValValLeuMetValLeu
   332. .554
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308. .836
== "probable primer binding site (primer #270)" . 825
  "tag= ]
note= "probable primer binding site (primer #271)"
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877. .893
   /*tag= m
/note= "probable primer binding site (primer #232)"
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product= "IA beta chain beta 2 region"
21. .550
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Matches:
Conservative:
Mismatches:
Indels:
   (NAJE-) NAT JEWISH CENT IMMUNOLOGY & RESPIRATORY
  [0-048-116B-6 (1-306) x AAT04262 (1-893)
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  94US-00207481
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      note=
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P-PSDB; AAR82533.
  cent Similarity:
: Local Similarity:
   WO9523814-A1.
   03-MAR-1995;
  04-MAR-1994;
  nment Scores:
  primer_bind
                        mat_peptide
   primer_bind
   primer_bind
  primer_bind
  08-SEP-1995
  Kappler JW,
```

SerSerProGlyThrGluGlyGlyAsnSerIleCysPheSerProSerLeuGluHisPro

```
121 TyrTrpasnSerGlnProGluileLeuGluArgThrArgAlaGluValAspThrAlaCys 140
  765
   226 GGTGGAGGCTCCGAAAGGCATTTCGTGGTCCAGTTCAAGGGCGAGTGCTACACCAAC 285
   100
   TyrAspSerAspValGlyGluTyrArgAlaValThrGluLeuGlyArgProAspAlaGlu 120
  465
  160
   525
  180
   200
  241 SerieulysSerProlleThrValGluTrpArgAlaGlnSerGluSerAlaArgSerLys 260
   I-Ab(beta)-Cholera toxin B subunit-leucine zipper (LZ)-BirA fusion cDNA.
  class II major histocompatibility complex; WHC; CD4+ T-cell detection; flow cytometry; mucous membrane invasive antigen; I-Ab(beta)-cholera toxin B subunit-leucine zipper-BirA fusion; CTB; 88;
  ValMetLeuGluMetThrProHisGlnGlyGluValTyrThrCysHisValGluHisPro
   706 GTCATGCTGGAGATGACCCCTCATCAGGGAGAGTCTACACCTGCCATGTGGAGGATCCC
                  286 GGGACGCAGCGCATACGGCTCGTGACCAGATACATCTACAACCGGGAGGAGGAGTACGTGCGC
   346 TACGACAGCGACGTGGGCGAGTACCGCGGGGGACCGAGCCGGGCGGCGCCAGACGCGGG
   406 TACTIGGAACAGCCAGCCGGAGATCCTGGAGCGAACGCGGGCCCAGGTGGACACGGCGTGC
   ValAlaileSerLeuSerArgThrGluAlaLeuAsnHisHisAsnThrLeuValCysSer
   ValThrAspPheTyrProAlaLysIleLysValArgTrpPheArgAsnGlyGluGlu
  61 GlyGlyGlySerGluArgHisPheValValGlnPheLysGlyGluCysTyrTyrThrAsn
   GlyThrGlnArglleArgLeuValThrArgTyrIleTyrAsnArgGluGluTyrValArg
   586 greacadarrrcracccadccaadarcaadrecccrecrrcagearreccadeade
   Thr ValGly ValSerSerThrGlnLeuIleArgAsnGlyAspTrpThrPheGlnValLeu
41 IleValValSerGlySerTrpAspGlyGlyGlyGlySerLeuValProArgGlySerGly
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  Location/Qualifiers
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   07-OCT-2004 (first entry)
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Unidentified.
   15-JUL-2004.
  161
  181
   201
  81
   101
   221
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  gene.
  RESULT 3
  ADQ31225
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(SENT-) SENTAN KAGAKU GIJUTSU INCUBATION CENT KK
03-DEC-2002; 2002JP-00351818.
```

WPI; 2004-546819/53. P-PSDB; ADQ31224.

Peptide-Class II major histocompatibility complex (WHC) composite, useful for detecting antigen specific CD4+ T-cell, comprises antigen peptide containing epitope of mucous membrane invasive protein, and extracellular region of MHC.

Example 1; SEQ ID NO 10; 30pp; Japanese

The invention relates to a novel class II major histocompatibility complex (MHC) antigenic peptide composite comprising a peptide containing the T-cell antigenic determinant of a mucous membrane invasive protein and the extracellular region of a class II MHC molecule or at least part of the extracellular region of the class II MHC molecule having an amino acid sequence comprising one or more deletions, substitutions or additions. The molecule of the invention may be useful for detecting an antigen-specific CD4+ T-cell by flow cytometry and for presenting a microorganism-derived mucous membrane invasive protein as an antigen. The method of the invention enables efficient detection of antigen-specific activation of CD4+ T-cells in the mucous membrane. The current sequence is that of the class II major histocompatibility complex-related I-cDNA of the invention.

Sequence 945 BP; 230 A; 256 C; 294 G; 165 T; 0 U; 0 Other;

| m     |
|-------|
| Gaps: |
| 12    |
|       |
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300
   124 GCGATCGCGGCCCATCAGCATGGCGAACGAGGTGGTGGTCC---GGTGGAGGGGGAAGT 180
   181 GGAGGTGGAGGGTCTGAAAGGCATTTCGTGTACCAGTTCATGGGCGAGTGCTACTTCACC 240
  ArgTyrAspSerAspValGlyGluTyrArgAlaValThrGluLeuGlyArgFroAspAla 119
   GluTyrTrpAsnSerGlnProGluIleLeuGluArgThrArgAlaGluValAspThrAla 139
  13
  59
   66
   LeuSerSerProGlyThrGluGlyGlyAsnSerIleCysPheSerProSerLeuGluHis 39
   79
   CGCTACGACAGCGACGTGGGGCGAGCACCGCGCGGTGACCGAGCTGGGGGCGGCCAGACGCC
   GlyGlyGlyGlySerGluArgHisPheValValGlnPheLysGlyGluCysTyrTyrThr
   CysArgHisAsnTyrGluGlyProGluThrSerThrSerLeuArgArgLeuGluGlnPro
   1 MetAlaLeuGlnIlePro---SerLeuLeuLeuSerAlaAlaValValValLeuMetVal
  40 ProlleValValSerGlySerTrpAspGlyGlyGlyGlySerLeuValProArgGlySer
   80 AsnGlyThrGlnArglleArgLeuValThrArgTyrIleTyrAsnArgGluGluTyrVal
3-048-116B-6 (1-306) x ADQ31225 (1-945)
  100
   9
   241
  301
  120
   361
   140
```

480

Tecagacacaactacgaegeccegagacccacacccrcccrecegecgcrrgaacagccc

```
Peptide-Class II major histocompatibility complex (MHC) composite, useful for detecting antigen specific CD4+ T-cell, comprises antigen peptide containing epitope of mucous membrane invasive protein, and extracellular
  199
  219
   600
   99
  239
   ProSerLeuLygSerProlleThrValGluTrpArgAlaGlnSerGluSerAlaArgSer 259
  280 LeulyslyslyslyslandalnLeulyslyplysleuGlnAlaLeulyslyslysleuAla 299
   1.915
/*taga -
/*taga -
//*taga -
//
   class II major histocompatibility complex; MHC; CD4+ T-cell detection; flow cytometry; mucous membrane invasive antigen; I-Ab(beta)-heat-labile toxin B subunit-leucine zipper-BirA fusion; LTB;
  ServalThrAspPheTyrProAlaLysIleLysValArgTrpPheArgAsnGlyGlnGlu
  GAGACGGTGGGGGTCTCATCCACACACACACTTATAGGAATGGGGACTGGACCTTCCAGGTC
AsnvalalalleSerLeuSerArgThrGluAlaLeuAsnHisHisAsnThrLeuValCys
                       LeuValMetLeuGluMetThrProHisGlnGlyGluValTyrThrCysHisValGluHis
  GluThrValGlyValSerSerThrGlnLeuIleArgAsnGlyAspTrpThrPheGlnVal
   LysGlyGlyGlyGlySerThrThrAlaProSerAlaGlnLeuLysLysLeuGlnAla
   I-Ab(beta)-E. coli heat-labile toxin B subunit-LZ-BirA fusion cDNA.
  ă
  (SENT-) SENTAN KAGAKU GIJUTSU INCUBATION CENT
   Location/Qualifiers
  BP.
  03-DEC-2003; 2003JP-00404367.
  03-DEC-2002; 2002JP-00351818.
  ADQ31228 standard; cDNA; 915
  07-OCT-2004 (first entry)
  300 GlnHisHisHis 303
   WPI; 2004-546819/53.
P-PSDB; ADQ31227.
  Escherichia coli.
Unidentified.
  JP2004196789-A.
  15-JUL-2004
   240
  260
   ADQ31228;
     160
   180
  200
   220
   661
  895
   601
  ss; gene
   RESULT 4
     8 8
  셤
   ઠે
   g
   g
  셤
   q
   ò
  g
  ò
   ઠે
   셤
   ઠે
   8
   ò
```

Ę region of Example 3; SEQ ID NO 13; 30pp; Japanese.

The invention relates to a novel class II major histocompatibility complex (MHC) antiganic peptide composite comprising a peptide containing the T-cell antigenic determinant of a mucous membrane invasive protein and the extracellular region of a class II MHC molecule or at least part of the extracellular region of the class II MHC molecule having an amino acid sequence comprising one or more deletions, substitutions or antigen-specific CD4+ T-cell by flow cytometry and for presenting an antigen-specific CD4+ T-cell by flow cytometry and for presenting a microorganism-derived mucous membrane invasive protein as an antigen. The method of the invention enables efficient detection of antigen-specific activation of CD4+ T-cells in the mucous membrane. The current sequence is that of the class II major histocompatibility complex-related I-bi(alpha)-Escherichia coli heat-labile toxin B subunit (ITB)-leucine zipper (LZ)-BirA fusion cDNA of the invention.

Sequence 915 BP; 228 A; 242 C; 271 G; 174 T; 0 U; 0 Other;

|                | 915       | 242     | 16              | 32                  | 14        | Ľ  |
|----------------|-----------|---------|-----------------|---------------------|-----------|----|
|                |           |         |                 | Mismatches:         |           |    |
|                | 7.14e-106 | 1219.00 | 84.9%           | 79.68               | 75.2\$    | 13 |
| inment Scores: | i. No.:   | .e:     | ent Similarity: | : Local Similarity: | :y Match: |    |

.0-048-116B-6 (1-306) x ADQ31228 (1-915)

- 19 63 1 MetAlaLeuGlnIlePro---SerLeuLeuLeuSerAlaAlaValValLeuMetVal
  - 20 LeuSerSerProGlyThrGluGlyGlyAsnSerIleCysPheSerProSerLeuGluHis 39
    - 64 creascrecearrescresersearcres 108
- 156 109 ATGGTTATCATT-----ACATTTAAGGGTGGTGGTGGTTCTTTAGTTCCTAGA-----40 ProllevalValSerGlySerTrpAspGlyGlyGlyGlySerLeuValProArgGlySer
- 157 -----GGTGGTAGTGAAAGGCATTTCGTGTACCAGTTCATGGGCGAGTGCTACTTCACC 210 79 GlyGlyGlyGlySerGluArgHisPheValValGlnPheLysGlyGluCysTyrThr 9
- 211 AACGGGACGCAGGGGATACGATATGTGACCAGATACATCTACAACCGGGAGGAGTACGTG 270 66 80 AsnGlyThrGlnArglleArgLeuValThrArgTyrIleTyrAsnArgGluGluTyrVal
- 100 ArgTyrAspSerAspValGlyGluTyrArgAlaValThrGluLeuGlyArgProAspAla 119
- 120 GluTyrTrpAsnSerGlnProGluIleLeuGluArgThrArgAlaGluValAspThrAla 139
- 140 CysArgHisAsnTyrGluGlyProGluThrSerThrSerLeuArgArgLeuGluGlnPro 159
- 160 AsnValAlaIleSerLeuSerArgThrGluAlaLeuAsnHisHisAsnThrLeuValCys 179 510
- SerValThrAspPheTyrProAlaLysIleLysValArgTrpPheArgAsnGlyGlnGlu 199 511 rchárdachdartricthcccháccchadhtchahárdcachdartccadhardacchdahá 570 180
- 219 GluThrvalGlyvalSerSerThrGlnLeuIleArgAsnGlyAspTrpThrPheGlnVal 200

Major histocompatibility class II antigen; MHC class II; T cell; T lymphocyte; Thi; Th2; activation; CD4+; antigen presenting cell; APC; autoimmune disease; diabetes; multiple sclerosis; autoimmune thyroiditis; systemic lupus erythromatosus; mysathenia gravis; Crohn's disease; inflammatory bowel disease; allery; asthma; contact sensitivity; immunotherapy; therapy; IAd beta chain; mouse; ds; circular; cyclic. 259 299 744 864 GAGACGGTGGGGGTCTCCACACACATATTAGGAATGGGGACTGGACCTTCCAGGTC 630 ProSerLeuLysSerProlleThrValGluTrpArgAlaGlnSerGluSerAlaArgSer LeuValMetLeuGluMetThrProHisGlnGlyGluValTyrThrCysHisValGluHis CCCAGCCTGAAGAGCCCCCATCACTGTGGAGTGGAGGGCACAGTCGTCAGCA-----GAC LeulyslyslyslysasnalaGlnLeulysTrpLysLeuGlnalaLeulysLysLysLysLeuAla LysGlyGlyGlyGlySerThrThrAlaProSerAlaGlnLeuLysLysLysLeuGlnAla CTGGTTCCGCGCGGATCCACTACAGCTCCATCAGCTCAGATGAAAAGAAACTGCAGGCA crtaagaaaagaacgcrcagcrgaagrggaactrcaagcccrcaagaagaaacrcgcc BP. AAV12068 standard; cDNA; 4724 Murine IAd beta chain cDNA. 08-JUN-1998 (first entry) GlnHisHisHis 303 CAGCTGCATCAT 876 AAV12068; 631 240 691 260 745 805 300 865 220 AAV12068 RESULT ઠે g ò 셤 ઠે 셤 ઠે 셤 ş 셤

97WO-US008697. 96US-0018175P. (SCRI ) SCRIPPS RES INST. W09746256-A1. 22-MAY-1997; 23-MAY-1996; 11-DEC-1997. Mus 

useful Synthetic antigen presenting cell for activating CD4+ T cells - use treat autoimmune disease, e.g. diabetes, multiple sclerosis, Crohn'disease and inflammatory bowel disease, or allergy, e.g. asthma and contact sensitivity. Peterson PA; Webb SR, Wingvist O, Karlsson L, Jackson MR, WPI; 1998-041895/04.

S

Example 2; Page 94-96; 141pp; English.

This nucleotide sequence comprises a PCR product obtained by amplification of mouse splenocyte cDNA using primers (see AAV12065 and AAV12066) designed for the amplification of IAd beta chain full-length cDNA. IAd alpha chain cDNA (see AAV12067) has been similarly obtained. The IAd sequences were cloned into metallothionein promoter (see AAV12062)-driven vector pRMHa-3 prior to sequencing. Major histocompatibility complex (MHC) class II IAd heterodimers were expressed at the cell surface of transfected Drosophila Schneider 2 (ATCC CRL 19974) cells. The invention relates to the preparation and use of synthetic antigen presenting matrices, in particular antigen presenting cells such as insect cells that have been transfected to produce MHC

||| ||||| ATGTTGAGCGCCATCGGGGGC 1149

1129

```
antigen presenting molecules with one or more accessory molecules. The matrices are used to activate naive CD4+ T cells and to shift the ongoing activation state into a preferred differentiated population of Th1 or Th2 cells. Applications include the treatment of autoimmune disease, e.g. diabetes, multiple sclerosis, autoimmune thyroiditis, systemic lupus erythematosus, myasthenia gravis, Crohn's disease and inflammatory bowel disease, or an allergy, e.g. asthma and contact sensitivity
```

Sequence 4724 BP; 1196 A; 1194 C; 1200 G; 1134 T; 0 U; 0 Other;

| 4724     | 228              | 7                                    | 1                               | 37                                                                                        | 7                                                                                 |
|----------|------------------|--------------------------------------|---------------------------------|-------------------------------------------------------------------------------------------|-----------------------------------------------------------------------------------|
| Length:  | Matches:         | Conservative:                        | Mismatches:                     | Indels:                                                                                   | Gaps:                                                                             |
| 1.85e-99 | 1161.50          | 85.8%                                | 85.48                           | 71.7                                                                                      | ~                                                                                 |
| . No     |                  | ant Similarity:                      | Local Similarity:               | / Match:                                                                                  |                                                                                   |
|          | 1.85e-99 Length: | 1.85e-99 Length:<br>1161.50 Matches: | 1.85e-99<br>1161.50<br>7: 85.8% | 1.85e-99 Length:<br>1161.50 Matches:<br>7: 85.8\$ Conservative:<br>1ty: 85.4\$ Mamatches: | 1.85e-99 Length:<br>1161.50 Matches:<br>7: 85.8\$ Conservative:<br>71.7\$ Indels: |

## 3-048-116B-6 (1-306) x AAV12068 (1-4724)

```
1128
  1008
   1068
  120
   140
  160
  240
   260
  708
   768
  828
   180
   888
  200
  588
   100
   648
  537
   537
20
   40
  9
  80
   MetAlaLeuGlnIleProSerLeuLeuLeuSerAlaAlaValValValLeuMetValLeu
                   SerSerProGlyThrGluGlyGlyAsnSerIleCysPheSerProSerLeuGluHisPro
   TACGACAGCGACGTGGGCGAGTACCGCGGGTGACCGAGCTGGGGCGGCGGCCAAG
  ACAGTGGGGGTCTCATCCACACAGCTTATTAGGAATGGGGACTGGACCTTCCAGGTCCTG
   GTCATGCTGGAGATGACCCCTCATCAGGAGAGGTCTACACCTGCCCATGTGGAGCATCCC
  41 IleValValSerGlySerTrpAspGlyGlyGlyGlySerLeuValProArgGlySerGly
  GlyGlyGlySerGluArgHisPheValValGlnPheLysGlyGluCysTyrThrAsn
  -----receaaagecarrregregrecaerreaagececaargererae
   GlyThrGlnArgIeArgLeuValThrArgTyrIleTyrAsnArgGluGluTyrValArg
   GGGACGCAGGCATACGGCTCGTGACCAGATACATCTACAACCGGGAGGAGTACGTGCGC
  TyrAspSerAspValGlyGluTyrArgAlaValThrGluLeuGlyArgProAspAlaGlu
   TyrTrpAsnSerGlnProGluIleLeuGluArgThrArgAlaGluValAspThrAlaCys
   ArgHisAsnTyrGluGlyProGluThrSerThrSerLeuArgArgLeuGluGlnProAsn
  AGACACAACTACGAGGGGCCGGAGACCAGCACCTCCCTGCGGCGGCTTGAACAGCCCAAT
   ValAlaileSerLeuSerArgThrGluAlaLeuAsnHisHisAsnThrLeuValCysSer
   ATCGCCATCTCCCTGTCCAGGACAGAGCCCCTCAACCACCACAAAAACACTCTGGTCTGTTCG
  ValThrAspPheTyrProAlaLysIleLysValArgTrpPheArgAsnGlyGluGlu
   GTGACAGATTTCTACCCAGCCAAGATCAAAGTGCGCTGGTTCAGGAATGGCCAGGAGGAG
   ThrvalGlyvalSerSerThrGlnLeuIleArgAsnGlyAspTrpThrPheGlnValLeu
  ValMetLeuGluMetThrProHisGlnGlyGluValTyrThrCysHisValGluHisPro
  1009
  241
-
                                 451
  21
  61
  538
  589
  101
   649
  709
  141
   769
   161
  829
  181
   889
  201
  949
   221
   537
   81
   121
```

------GlyGlyGlyGly 264

261

```
/*tag= b
note= "binding site for primer #362 (see AAT04270)"
   (see AAT04271)"
   /*tag= h
/note= "probable primer binding site (primer #270)"
31. .959
  /*tag= k
note= "probable primer binding site (primer #272)"
323. .850
  '*tag= j
'note= "probable primer binding site (primer #271)"
544. .568
  cell;
   primer binding site (primer #259)"
  (primer #366)"
  /*tag= a
/note= "probable primer binding site (primer #76)"
complement(40. .74)
  #29) "
   /*tag= g
note= "primer #365 (see AAT04272) binding site"
  Major histocompatability complex; MHC; T-cell receptor; TCR; autoimmune disease; immunodeficiency disease; immune response; immunoproliferation disease; graft-host rejection; therapy; B
  /note= "probable primer binding site (primer
  "IA beta chain beta 2 region"
   /*tag= e
/note= "binding site for primer #363
complement(177. .226)
   /*tag= m
/note= "probable primer binding site
(000. .1013
  gite
   (NAJE-) NAT JEWISH CENT IMMUNOLOGY & RESPIRATORY
  chain"
  /*tag= f
/note= "primer #364 binding
complement(212, 266)
   /*tag= c
/product= "hybrid IA beta
   /*tag= d
/note= "leader region"
complement(140. .191)
  Location/Qualifiers
   /*tag= 1
/note= "probable p
942. .976
          BP.
          1013
   94US-00207481
   95WO-US002689
   product=
  (first entry)
  403
   Hybrid IA beta chain gene
   535. .564
   63. .959
   63. .143
           DNA;
   M12.C3; pM12-IAb-Ea; 88.
   ..18
*tag=
   *tag=
          standard;
   WO9523814-A1
  03-MAR-1995;
  16-APR-1996
   primer_bind
  primer_bind
   38-SEP-1995
  primer_bind
  primer_bind
  sig_peptide
  primer_bind
  primer_bind
  primer_bind
  mat_peptide
   primer_bind
  primer_bind
   primer_bind
   primer_bind
   Synthetic
                              AAT04269;
           AAT04269
```

```
ä
Marrack
                       WPI; 1995-320543/41.
                                 P-PSDB, AAR82538
Kappler JW,
```

Peptide-MHC complex comprising antigenic peptide, linker and MHC segment - useful as reagents for the treatment of diseases including auto-immune diseases, immuno-stimulatory diseases or graft-host rejection.

Example 2; Page 65; 94pp; English.

This sequence represents a hybrid IA beta chain gene. This sequence contains a fragment of the IB alpha chain (residues 56-73), as well as a linker and cleavage site. This sequence was transfected into a B cell line (M12.C3) using plasmid pM12-IAb-Ba. It was found that the encoded sequence was expressed in these cells. Complexes such as this may be used to regulate an immune response. The complexes are capable of being trecognised by a TCR alone or in combination with additional MHC proteins. These complexes are useful for therapeutic purposes and experimental purposes. They can also be used as reagents for the treatment of diseases including autolumnume diseases, immunodeficiency diseases, immunoproliferation diseases, and graft-host rejection

Sequence 1013 BP; 220 A; 272 C; 327 G; 192 T; 0 U; 2 Other;

| 1013     | 230              | 9                                    | 22                                                          | 14                                                                               | 4                                                                                                                                                                                 |
|----------|------------------|--------------------------------------|-------------------------------------------------------------|----------------------------------------------------------------------------------|-----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| Length:  | Matches:         | Conservative:                        | Mismatches:                                                 | Indels:                                                                          | Gana:                                                                                                                                                                             |
| 2.31e-99 | 1151.00          | 86.8%                                | 84.6%                                                       | 71.0\$                                                                           | 0                                                                                                                                                                                 |
| i. No.:  | .e.:             | sent Similarity:                     | : Local Similarity:                                         | .y Match:                                                                        |                                                                                                                                                                                   |
|          | 2.31e-99 Length: | 2.31e-99 Length:<br>1151.00 Matches: | 2.31e-99 Length:<br>1151.00 Matches:<br>86.8% Conservative: | 2.31e-99 Length:<br>1151.00 Matches:<br>86.84 Conservative:<br>84.68 Mismatches: | i. No.: 2.31e-99 Length: 1013<br>:en: 1151.00 Matches: 230<br>:ent Similarity: 86.8\$ Conservative: 6<br>: Local Similarity: 84.6\$ Mismatches: 22<br>:y Match: 71.0\$ Indels: 14 |

|                                                                                |                                         | 20                                                          | 40                                                              | 55  | 75<br>284                                                           | 95<br>344                                                                     | 115                                                                 | 135                                                                 | 155<br>524                                                           | 175<br>584                                                       | 195                                                             |
|--------------------------------------------------------------------------------|-----------------------------------------|-------------------------------------------------------------|-----------------------------------------------------------------|-----|---------------------------------------------------------------------|-------------------------------------------------------------------------------|---------------------------------------------------------------------|---------------------------------------------------------------------|----------------------------------------------------------------------|------------------------------------------------------------------|-----------------------------------------------------------------|
| y: 66.8% CONBELVALIVE: 6 Tity: 84.6% Mismatches: 22 71.0% Indels: 14 2 Gaps: 4 | -048-116B-6 (1-306) x AAT04269 (1-1013) | 1 MetAlaLeuGhnIleProSerLeuLeuSerAlaAlaValValValLeuMetValLeu | SerSerProGlyThrGluGlyGlyAsnSerIleCysPheSerProSerLeuGluHisPro 40 | 41  | ProArgGlySerGlyGlyGlyGlySerGluArgHisPheValValGlnPheLysGlyGlu 75<br> | CystyrtyrthrasnGlythrGlnarglleargLeuValthrargTyrIletyrasnarg 95<br>        :: | GluGluTyrvalargTyraspSeraspValGlyGluTyrargAlavalThrGlùLeuGly 11<br> | ArgProAspalaGluTyrTrpAsnSerGlnProGluIleLeuGluArgThrArgAlaGlu 13<br> | Valaspihralacysarghisassifyrdluglyprogluthrserthrserleuargarg 15<br> | LeugluginproasnyalalaileserLeuserargthrdlualaLeuasnHisHsn 17<br> | ThrLeuValCysSerValThrAspPheTyrProAlaLysIleLysValArgTrpPheArg 19 |
| m in                                                                           | 9-e                                     | A—A                                                         | 8—8<br>8—8                                                      | ; 8 |                                                                     | \$=\$                                                                         |                                                                     |                                                                     |                                                                      |                                                                  |                                                                 |
| nc Simitaricy:<br>Local Similarity:<br>Match:                                  | -048-116                                | 63                                                          | 21                                                              | 41  | 56<br>228                                                           | 76                                                                            | 96<br>345                                                           | 116                                                                 | 136                                                                  | 156<br>525                                                       | 176                                                             |

```
216 ThrPheGlnValLeuValMetLeuGluMetThrProHisGlnGlyGluValTyrThrCys 235
  236 HisValGluHisProSerLeuLysSerProlleThrValGluTrpArgAlaGlnSerGlu 255
   704
   764
   6. . 86
/*teg= b
hote= I-Ad beta chain leader
/note= "murine MHC clase II I-Ad gene beta chain leader
  MHC; major histocompatibility complex; PCR; polymerase chain reaction; T cell activity modulator; antagonist; immune disorder; allergy; multiple sclerosis; insulin-dependent diabetes mellitus; rheumatoid arthritis; myashenia gravis; ds.
196 AsnGlyGlnGluGluThrValGlyValSerSerThrGlnLeuIleArgAsnGlyAspTrp
  /*tag= h
/label= I-Ad_alphal
/note= "murine MHC class II I-Ad gene alpha-2 domain"
1068. .1352
   class II I-Ad gene alpha-2 domain"
   /*tag= e
/label= I-Ad betal
/label= "murine MHC class II I-Ad gene beta-1 domain"
453. 734
  /*tag= f
/label= I-Ad_beta2
/label= "murine MHC class II I-Ad gene beta-2 domain"
/35. .806
   Vector SCE1-derived single chain gene encoding MHC fusion complex.
  /*tage c
/label= OVA 323-339
/note= "chicken ovalbumin residues 323-339"
138. 167
   "rtag= g
'note= "24 residue peptide linker"
007. .1067
  /*tag= d
/note= "10 residue linker peptide"
   264
   825 rcreccreaaccaacaacarerraaccecarcecece 860
   SerAlaArgSerLys-----GlyGlyGlyGly
   /*tag= 1
/label= I-Ad alpha2
/note= "murine MHC cl
1353. .1379
  Location/Qualifiers
6. .1382
/*tag= a
  BP
  /*tag= j
/note= "EB tag"
  AAT17588 standard; DNA; 1382
  (first entry)
  sequence"
87. .137
   .452
   26-SEP-1996
  misc_feature
  misc_feature
   misc_feature
   misc_feature
  misc_feature
   misc_feature
   misc_feature
  misc_feature
  sig peptide
  Synthetic.
   AAT17588;
   256
   AAT17588
ID AAT1
  Key
   RESULT 7
   HILLIGHT CHARLES CONTROL CONTR
  요
  ઠે
   셤
   셤
  ઠે
  ઠે
  셤
   ઠે
```

```
27-MAR-1998 (first entry)
  6. .1382
/*tag= a
   SCE1 single chain gene.
  (DADE-) DADE INT INC.
   WPI; 1997-402555/37.
P-PSDB; AAW29214.
  30-JAN-1997;
  31-JAN-1996;
  07-AUG-1997.
  WO9728191-A1
   Synthetic.
  241
  261
  AAT86989;
   121
   201
101
   221
   8 6
   q
   g
   ð
  g
   g
   à
  셤
   g
  ሯ
   요
   ò
   셤
  δ
  ò
  ð
   ò
   AAT17588 encodes a murine MHC fusion complex capable of modulating T cell activity encoded by the vector SCEI. The MHC fusion complex comprises at least one MHC molecule containing a peptide-binding groove and a presenting peptide covalently linked to the MHC molecule and opt. a transmembrane domain. DNA encoding a MHC fusion complex may be cloned into a host cell to express the complex. The transformed cells may then be used to identify peptides that modulate, pref. antegonie, T cell activity. DNA encoding a MHC fusion complex or a single chain fusion molecule may be used to vaccinate a mammal against a targeted disorder. The fusion complexes may be used to suppress an immune response in an animal suffering from an immune disorder e.g. multiple sclerosis, insulin-dependent disbetes mellitus, rheumatoid arthritis myssthemia gravis or chronic allergies. The complexes may also be used in the treatment of livestock and pets such as cats and dogs. The MHC fusion complexes can be produced such that they contain a single antigenic peptide including one of known structure, additionally a wide range of peptides can be presented for T cell interaction
   120 ATCAACGAAGCTGGTCGTCGTAGCGGAAGGGGGCGGAAGC---------GGCGGA 164
   165 dedecanacticidana de cartificat de fica esticana de describilitados de describilidades de describilidades de describilidades de decembros de describilidades de decembros 
   6 ATGGCTCTGCAGATCCCCAGCCTCCTCTCAGCTGCTGGTGGTGGTGCTGATGGTGCTG 65
  9
   80
   21 SerSerProGlyThrGluGlyGlyAsnSerIleCysPheSerProSerLeuGluHisPro 40
  41 IleValValSerGlySerTrpAspGlyGlyGlyGlySerLeuValProArgGlySerGly
   61 GlyGlyGlyGlySerGluArgHisPheValValGlnPheLy8GlyGluCysTyrTyrThrAsn
  1 MetAlaLeuGlnIleProSerLeuLeuLeuSerAlaAlaValValLeuMetValLeu
   Major histocompatability complex fusion complex for modulating T ceactivity - used in the treatment of immune disorders, e.g. multiple sclerosis, IDDM and rheumatoid arthritis.
  Sequence 1382 BP; 320 A; 374 C; 404 G; 284 T; 0 U; 0 Other;
  Edwards AC;
  1382
227
4
24
10
3
  Length:
Matches:
Conservative:
Mismatches:
  ŝ
   Grammer
   3-048-116B-6 (1-306) x AAT17588 (1-1382)
  Example 17; Fig 29; 210pp; English
  Weidanz JA,
   95WO-US009816
   94US-00283302
95US-00382454
  1.3e-98
1145.00
87.2%
85.7%
   Wong HC, Rhode PR, W
Chavaillaz P, Jiao J;
   (DADE-) DADE INT INC.
   WPI; 1996-129343/13.
P-PSDB; AAR98907.
  ent Similarity:
Local Similarity:
     WO9604314-A1
   31-JUL-1995;
  29-JUL-1994;
  01-FEB-1995;
   ment Scores:
  15-PEB-1996.
   y Match:
```

```
464
  584
   644
   704
   Single chain major histocompatibility complex comprising linked alpha and
beta chains – useful for suppressing an immune response to an auto:immune
   404
   141 ArgHisAsnTyrGluGlyProGluThrSerThrSerLeuArgArgLeuGluGlnProAsn 160
  180
   524
  200
  220
  ValMetLeuGluMetThrProHisGlnGlyGluValTyrThrCysHisValGluHisPro 240
   SerLeuLysSerProlleThrValGluTrpArgAlaGlnSerGluSerAlaArgSerLys 260
   TyraspSerAspValGlyGluTyrargAlavalThrGluLeuGlyArgProAspAlaGlu
               TyrTrpAsnSerGlnProGluIleLeuGluArgThrArgAlaGluValAspThrAlaCys
   ThrValGlyValSerSerThrGlnLeuIleArgAsnGlyAspTrpThrPheGlnValLeu
  construction, major histocompatibility complex; MHC; fusion complex; SCE1 single chain gene; se.
   HC;
  Wong
   Location/Qualifiers
   Rhode PR, Jiao J, Burkhardt M,
   86
   AAT86989 standard; DNA; 1382
   97WO-US001617.
  96US-00596387
   Glyglyglyglyser 265
   756 ecceditericarice 770
```

770

```
120
  140
   180
   200
   220
  644
   ValMetLeuGlumetThrProHisGlnGlyGluValTyrThrCysHisValGluHisPro 240
   755
   66 AGCAGCCCAAGGACCTTAAGTATCTCTCAGGCTGTTCACGCTGCTCACGCTGAA----- 119
  100
   284
   344
   404
  ArgHisAsnTyrGluGlyProGluThrSerThrSerLeuArgArgLeuGluGlnProAsn 160
  464
  524
  584
  164
  224
   21 SerSerProGlyThrGluGlyGlyAsnSerIleCysPheSerProSerLeuGluHisPro 40
   9
   8
  165 GGGGGAAACTCCGAAAGGCATTTCGTGGTCCAGTTCAAGGGCGAGTGCTACTACAAC
   GlyThrGlnArglleArgLeuValThrArgTyrIleTyrAsnArgGluGluTyrValArg
   225 GGACGCAGCGCATACGCCTCGTGACCAGATACATCTACAACCGGGAGGAGTACGTGCGC
  101 TyraspSeraspValGlyGluTyrArgAlaValThrGluLeuGlyArgProAspAlaGlu
   285 raceacadedaceredeceaeracedecedereaeceaacredecedeceaecedad
   345 TACTGGAACAGCCAGCCGGAGATCCTGGAGCGAACGCGGGCCGAGGTGGACACGCGTGC
  AGACACAACTACGAGGGGCCGGAGACCAGCACCTCCCTGCGGCGCGTTGAACAGCCCAAT
   ValAlaIleSerLeuSerArgThrGluAlaLeuAsnHisHisAsnThrLeuValCysSer
   GTCGCCATCTCCCTGTCCAGGACAGAGGCCCTCAACCACCACAAAAACACTCTGGTCTGTTCG
   ValThrAspPheTyrProAlaLyslleLysValArgTrpPheArgAsnGlyGluGlu
   GTGACAGATTTCTACCCAGCCAAGATCAAAGTGCGCTGGTTCAGGAATGGCCAGGAGGAG
   ThrvalGlyValSerSerThrGlnLeuIleArgAsnGlyAspTrpThrPheGlnValLeu
   SerieulysSerProlleThrValGluTrpArgAlaGlnSerGluSerAlaArgSerLys
   705 AGCCTGAAGAGCCCCATCACTGTGGAGTGG-----ACTAGTGGTGGCGGTGGCAGC
   MetAlaLeuGlnIleProSerLeuLeuLeuSerAlaAlaValValValLeuMetValLeu
  120 ATCAACGAAGCTGGTCGTGCTAGCGGAGGGGGGGGGAAGC-------GGCGGA
  TyrTrpAsnSerGlnProGluIleLeuGluArgThrArgAlaGluValAspThrAlaCys
   41 ilevalvalSerGlySerTrpAspGlyGlyGlyGlySerLeuvalProArgGlySerGly
disease, e.g. multiple sclerosis, rheumatoid arthritis, diabetes
  Sequence 1382 BP; 320 A; 373 C; 405 G; 284 T; 0 U; 0 Other;
   of major
   the construction
   fusion complexes
  Length:
Matches:
Conservative:
Mismatches:
Indels:
  Example 17, Page 140-141; 217pp; English.
   Gaps:
  .0-048-116B-6 (1-306) x AAT86989 (1-1382)
   The present sequence was used in histocompatibility complex (MHC)
  261 GlyGlyGlyGlySer 265
   1.3e-98
1145.00
87.2%
85.7%
  cent Similarity:
Local Similarity:
y Match:
                       mellitus, etc.
   nment Scores:
  141
  405
   191
  465
   181
   201
  221
   645
  241
  121
  81
```

```
The invention relates to a major histocompatibility complex (WHC) fusion complex (I) comprising an WHC molecule that contains a peptide-binding groove, and a presenting peptide covalently (e.g. an antigenic peptide) linked to the MHC molecule, where (I) is capable of modulating the cativity of a T cell. Also included are a DNA construct coding for the activity of a T cell. Also included are a DNA construct coding for the complex, where the MHC molecule is a class II MHC (e.g. mouse I-Ad or I-As, or human HLA-DR1 (human leukocyte antigen-DR1), a wultivalent MHC tusion complex comprising vectors that each contain the fusion complex. DNA, cluturing the host cells under conditions sultable for expression of the culturing the host cells under conditions sultable for expression of the Complex, and selecting host cells that express MHC fusion complex, and selecting host cells that axpress MHC fusion complex, and selecting host cells that axpress MHC fusion of the fusion complex MHC process, a single recombinant captures that modulate the activity of T cells), a single recombinant captures and beta chains of the MHC fusion complex. The DNA constructs and the alpha and beta chains of the fusion complex. Also included are inducing an immune complex, and sequence coding for a fusion complex. Also included are inducing an immune cresponse in a mammal including vaccinating a single chain fusion complex. DNA sequence coding for a fusion complex which is a single chain fusion complex. Or DNA sequence coding for a fusion complex which is a single chain fusion complex or encounting and a mammal and a mammal and a mammal and mumulant complex which is a single chain fusion complex or compression weeter, encoding a full length MHC molecule that contains a transmembrane domain, and a contains a transmembrane domain, and a contains a transmembrane domain, and a cell is presenting peptide that is a T cell receptor (TCR) antagonist or partial
   Novel major histocompatibility complex fusion complex having presenting peptide covalently linked to MHC molecule containing peptide-binding groove, used for suppressing immune response in multiple sclerosis,
   MHC; major histocompatibility complex; gene therapy; fusion complex; peptide-binding groove; T cell modulation; class II MHC; vaccine; autorimmune disorder; multiple sclerosis; rheumatoid arthritis; insulin-dependent diabetes mellitus; myasthenia gravis; immunogen; chronic allergy; mouse; ds; I-Ad; gene.
   Grammer S, Edwards AC;
   Mouse MHC I-Ad/Ova 323-339 synthetic gene SCE1.
  Example 17; Fig 29; 126pp; English.
   Weidanz JA,
BP.
   94US-00283302.
95US-00382454.
97US-00776084.
DNA; 1382
  06-JUL-2001; 2001US-00900379
   (first entry)
   Chavaillaz P, Jiao JJJ;
  (DADE-) DADE INT INC.
   Rhode PR,
   WPI; 2003-341126/32.
ACA60744 standard;
  P-PSDB; ABU72108
   US2002198144-A1
   29-JUL-1994;
01-FEB-1995;
17-JAN-1997;
   16-JUN-2003
  26-DEC-2002
   Mus sp.
Synthetic.
  allergies.
  ACA60744;
   HC,
  Wong
```

```
agonist and is covalently linked to the MHC protein, or DNA sequence coding for the fusion complex which is a single chain fusion molecule. The methods are useful for identifying a peptide that can modulate the activity of T cells, inducing an immune response in a mammal (including vaccinating a mammal against a targeted disorder) and for suppressing an immune response in a mammal. The disorders include an autoimmune disorder such as multiple sclerosis, insulin-dependent disbetes mellitus, rheumatoid arthritis, myasthenia gravis or chronic allergies. The present sequence encodes a mouse MHC class II I-Ad fusion complex of the
```

Sequence 1382 BP; 320 A; 373 C; 405 G; 284 T; 0 U; 0 Other;

```
120
   344
  140
   160
   200
   SerLeuLy8SerProlleThrValGluTrpArgAlaGlnSerGluSerAlaArgSerLy8 260
   100
   284
   404
  464
   ValalalleSerLeuSerArgThrGluAlaLeuAsnHisHisAsnThrLeuValCysSer 180
  524
   584
   644
  704
   164
  GGGGGAAACTCCGAAAGGCATTTCGTGGTCCAGTTCAAGGGCGAGTGCTACTACACCAAC 224
  9
   80
   20
   65
  SerSerProGlyThrGluGlyGlyAsnSerIleCysPheSerProSerLeuGluHisPro 40
   GTCATGCTGGAGATGACCCCTCATCAGGAGAGGTCTACACCTGCCATGTGGAGCATCCC
   AFGGCTCTGCAGATCCCCAGCCTCCTCTCAGCTGCTGTGGTGGTGGTGTTGATGGTGCTGT
  GlyThrGlnArgIleArgLeuValThrArgTyrIleTyrAsnArgGluGluTyrValArg
  GGGACGCAGCGCATACGGCTCGTGACCAGATACATCTACAACCGGGAGGAGTACGTGGC
  TyrAspSerAspValGlyGluTyrArgAlaValThrGluLeuGlyArgProAspAlaGlu
   ValThrAspPheTyrProAlaLysIleLysValArgTrpPheArgAsnGlyGluGlu
  ThrValGlyValSerSerThrGlnLeulleArgAsnGlyAspTrpThrPheGlnValLeu
   MetAlaLeuGlnIleProSerLeuLeuLeuSerAlaAlaValValValLeuMetValLeu
  41 IleValValSerGlySerTrpAspGlyGlyGlyGlySerLeuValProArgGlySerGly
   120 ATCAACGAAGCTGGTCGTGCTAGCGGAGGGGGGGGAAGC------GGCGGA
   GlyGlyGlyGlySerGluArgHisPheValValGlnPheLysGlyGluCysTyrTyrThrAsn
  TyrTrpAsnSerGlnProGluIleLeuGluArgThrArgAlaGluValAspThrAlaCys
   ArgHisAsnTyrGluGlyProGluThrSerThrSerLeuArgArgLeuGluGlnProAsn
   Greacagartrictacceageceaagarcaaagrececregircaggaareceaggageag
   ACAGTGGGGGTCTCATCCACACAGCTTATTAGGAATGGGGACTGGACCTTCCAGGTCCTG
  ValMetLeuGluMetThrProHisGlnGlyGluValTyrThrCysHisValGluHisPro
               Length:
Matches:
Conservative:
Mismatches:
Indels:
   Gaps:
  (1-1382)
  )-048-116B-6 (1-306) x ACA60744
               1.3e-98
1145.00
87.2
85.7
70.7
  ant Similarity:
Local Similarity:
ment Scores:
  21
   61
  165
   81
  225
   101
   285
  121
   345
  141
  405
   161
  465
   181
  525
   201
   585
  221
   645
  241
```

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--ACTAGTGGTGGCGGTGGCAGC 755
   6. .86
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/label= I-Ad_beta_chain_leader
/loce= "murine MHC class II I-Ad gene beta chain leader
sequence"
  WHC; major histocompatibility complex; PCR; polymerase chain reaction; T cell activity modulator; antagonist; immune disorder; allergy; multiple sclerosis; insulin-dependent diabetes mellitus; rheumatoid arthritis; myasthenia gravis; ds.
   gene alpha-2 domain"
   domain"
  domain"
  domain"
  encoding MHC fusion complex
   gene alpha-2
  beta-2
  gene beta-1
   87. .137
/ttag= c
/label= OVA_323-339
/note= "chicken ovalbumin residues 323-339"
138. .167
   Edwards AC;
  gene
  /*tag= d
/note= "10 residue linker peptide"
168. 452
  /*tag= g
/note= "24 residue peptide linker"
107. .1067
  class II I-Ad
   /*tag= e
/label= I-Ad_betal
/note= "murine WHC class II I-Ad
453. .734
  class II I-Ad
  II I-Ad
   Grammer S,
  /*tag= h
/label= I-Ad_alphal
/note= "murine MHC class
1068. .1352
AGCCTGAAGAGCCCCATCACTGTGGAGTGG-
  Vector SSC1-derived single chain gene
   /*tag= i
/label= I-Ad_alpha2
/note= "murine MHC c.
1353. 1382
   /*tag= f
/label= I-Ad_beta2
/note= "murine MHC c
735. .806
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6. .1385
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   Rhode PR, Weidanz JA,
  BP.
   "6xH18
  AAT17586 standard; DNA; 1385
  95WO-US009816
   94US-00283302
95US-00382454
   (first entry)
                   261 GlyGlyGlyGlySer
                                    GCGGTGGTGGTTCC
  (DADE-) DADE INT INC
   29-JUL-1994;
01-FEB-1995;
   misc_feature
  misc_feature
  31-JUL-1995;
  misc_feature
  misc_feature
  WO9604314-A1
   26-SEP-1996
   misc_feature
   misc_feature
   misc_feature
  misc feature
  15-FEB-1996
  sig_peptide
   Synthetic
  AAT17586;
 705
                                     756
   Wong HC,
  RESULT 10
   Key
   Š
```

```
Chavaillaz P, Jiao J;

WPI; 1996-129343/13.
P-PSDB; AAR98905.
Major histocompatability complex fusion complex for modulating T cell activity - used in the treatment of immune disorders, e.g. multiple sclerosis, IDDM and rheumatoid arthritis.

Example 17; Fig 27; 210pp; English.

AAT17586 encodes a murine MHC fusion complex capable of modulating T cell activity encoded by the vector SSC1. The MHC fusion complex comprises at least one MHC molecule containing a peptide-binding groove and a presenting peptide covalently linked to the MHC molecule and opt. a transmembrane domain. DNA encoding a MHC fusion complex may be cloned into a host cell to express the complex. The transformed cells may then be used to identify peptides that modulate, pref. antagonise, T cell activity. DNA encoding a MHC fusion complex or a single chain fusion molecule may be used to vaccinate a mammal against a trargeted disorder. The fusion complexes may be used to suppress an immune response in an animal suffering from an immune disorder e.g. multiple sclerosis, insulindependent diabetes may be used to suppress an immune response in an animal suffering from an immune disorder e.g. multiple sclerosis, insulindependent diabetes may also be used in the treatment of thromic allergies. The complexes may also be used in the treatment of thromic allergies. The complexes may also be used in the treatment of produced such that they contain a single antigenic peptide including one. Of known structure, additionally a wide range of peptides can be presented for T cell interaction
```

Sequence 1385 BP; 316 A; 384 C; 398 G; 287 T; 0 U; 0 Other;

|                                                                                   |                                  | ValLeuMetValLeu 20<br>                                           | oSerLeuGluHisPro 40<br>   <br>TCACGCTGAA 119                 | lProArgGlySerGly 60<br>   <br>GGCGGA 164                      | uCysTyrThrasn 80<br>                                             | gGluGluTyrValArg 100<br>                                      | yArgProAspAlaGlu 120<br>                                         | uvalaspThralaCys 140<br>                                        | gLeuGluGlnProAsn 160<br>                                          |
|-----------------------------------------------------------------------------------|----------------------------------|------------------------------------------------------------------|--------------------------------------------------------------|---------------------------------------------------------------|------------------------------------------------------------------|---------------------------------------------------------------|------------------------------------------------------------------|-----------------------------------------------------------------|-------------------------------------------------------------------|
| 1385<br>227<br>4<br>24<br>10<br>3                                                 |                                  | avalva<br>       <br>rgreer                                      | eSerPr<br>:::<br>:GCTGC                                      | rLeuVa<br> <br>                                               | 3G17G1<br>       <br>3GGCGA                                      | rAsnAr<br>       <br>caaccg                                   | uleug]<br>       <br>3CTCGG                                      | aAlagi<br>      <br>seccea                                      | uArgAr<br>       <br>scggcg                                       |
| Length: Matches: Conservative: Mismacches: Indels:                                | 6 (1-1385)                       | MetalaleuglnileProserleuleuleuseralaalavalvalvalleumetValleu<br> | SerSerProGlyThrGluGlyGlyAsnSerIleCysPheSerProSerLeuGluHisPro | IlevalvalSerGlySerTrpAspGlyGlyGlySerLeuvalProArgGlySerGly<br> | GlyGlyGlySerGlukrgHisPheValValGlnPheLygGlyGluCysfyrTyrThrAsn<br> | Glythrglinargileargleuvalthrargfyrilefyrasnarggluglutyrvalarg | TyrabpserabpValGlyGluTyrargalaValThrGluLeuGlyargFroaspAlaGlu<br> | Tyrtrpaenserginproglutieleugluargthrarglagluvalaepthralacye<br> | Arghi sasnTyrGluGlyProGluThrSerThrSerLeuArgArgLeuGluGlnProAsn<br> |
| 1.3e-98<br>1145.00<br>87.2%<br>85.7%<br>70.7%                                     | :0-048-116B-6 (1-306) x AAT17586 | IGINI LEProSe<br>            <br> CAGATCCCCAG                    | oglyThrGluGl;<br>   <br>AAGGACCTTAAG                         | SerGlySerTrj<br>:::   <br>\GCTGGTCGTGC                        | 'SerGluArgHi<br>          <br>TCCGAAAGGCA'                       | ArgileArgle<br>           <br>sccatacect                      | raspvalglygl<br>          <br> GACGTGGGCGA                       | nSerGlnProGl                                                    | TyrgluglyPr<br>          <br>TACGAGGGGCC                          |
| Scores:<br>milarity:<br>Similarity:<br>h:                                         | B-6 (1-306                       |                                                                  |                                                              |                                                               |                                                                  | -                                                             |                                                                  | -                                                               |                                                                   |
| <pre>jnment Scores: 1. No.: 7e: 2e: iont Similarity Local Similar ry Match:</pre> | :0-048-116                       | H 9                                                              | 21 66                                                        | 41                                                            | 61<br>165                                                        | 81 225                                                        | 101                                                              | 121                                                             | 141                                                               |

WHC; major histocompatibility complex; gene therapy; fusion comple peptide-binding groove; T cell modulation; class II MHC; vaccine; autoimmune disorder; multiple sclerosis; rheumatoid arthritis; insulin-dependent diabetes mellitus; myasthenia gravis; immunogen; chronic allergy; mouse; ds; I-Ad; gene.

Mus

```
180
  TyrAepSerAepValGlyGluTyrArgAlaValThrGluLeuGlyArgProAepAlaGlu 120
  140
  644
   240
   755
  224
   284
   344
   404
  524
  704
  SerLeuLysSerProlleThrValGluTrpArgAlaGlnSerGluSerAlaArgSerLys 260
  164
   9
   80
  65
   40
  66 AGCAGCCCAAGGACCTTAAGTATCTCTCAGGCTGTTCACGCTGCTCACGCTGAA-----
   GGGGGAAACTCCGAAAGGCATTTCGTGGTCCAGTTCAAGGCGAGTGCTACTACACCAAC
   GlyThrGlnArg1leArgLeuValThrArgTyrIleTyrAsnArgGluGluTyrValArg
   TyrTrpAsnSerGlnProGluIleLeuGluArgThrArgAlaGluValAspThrAlaCys
  ArgHisAsnTyrGluGlyProGluThrSerThrSerLeuArgArgLeuGluGlnProAsn
   ValalalleSerLeuSerArgThrGluAlaLeuAsnHisHisAsnThrLeuValCysSer
  ValThrAspPheTyrProAlaLysIleLysValArgTrpPheArgAsnGlyGluGlu
  GTGACAGATTTCTACCCAGCCAGCCAAGATCAAAGTGCGCTGGTTCAGGAATGGCCAGGAGAG
   ThrvalGlyvalSerSerThrGlnLeuIleArgAsnGlyAspTrpThrPheGlnValLeu
   ACAGTGGGGGTCTCATCCACACGCTTATTAGGAATGGGGACTGGACCTTCCAGGTCCTG
   ValMetLeuGluMetThrProHisGlnGlyGluValTyrThrCysHisValGluHisPro
   GTCATGCTGGAGATGATGACCCCTCATCAGGGAGGTCTACACCTGCCATGTGGAGGCATCCC
  MetAlaLeuGlnIleProSerLeuLeuLeuSerAlaAlaValValValLeuMetValLeu
  SerSerProGlyThrGluGlyGlyAsnSerIleCyaPheSerProSerLeuGluHiaPro
   IlevalvalSerGlySerTrpAspGlyGlyGlyGlySerLeuValProArgGlySerGly
  120 Archacgaagcregregretagceaaceaeaeaeaeaaac------Gecea
   GlyGlyGlySerGluArgHisPheValValGlnPheLysGlyGluCysTyrTyrThrAsn
   TACGACAGCGACGTGGGCGAGTACCGCGGGGGGAGCTGGGGGCGGCCAGACGCCGAG
   TACTIGGAACAGCCAGCCGGAGATCCTGGAGCGAACGCGGGCCGAGGTGGACACGGCGTGC
   AGACACAACTACGAGGGGCCGGAGACCAGCACCTCCCTGCGGCGGCGCTTGAACAGCCCAAT
   705 AGCCTGAAGAGCCCCATCACTGTGGAGTGG-----ACTAGTGGTGGCGGTGGCAGC
     3 2 4
Conservative:
Mismatches:
Indels:
  Gaps:
   J-048-116B-6 (1-306) x AAT86987 (1-1385)
   BP
  GlyGlyGlyGlySer 265
   ACA60742 standard; DNA; 1385
   (first entry)
 ant Similarity:
Local Similarity:
   16-JUN-2003
   21
   41
   61
  165
   81
  101
   285
  121
   345
  141
   405
   161
  465
   181
  525
   201
   221
  241
  261
   756
  ACA60742
                                       / Match:
```

```
The invention relates to a major histocompatibility complex (MHC) fusion complex (I) comprising an MHC molecule that contains a peptide-binding groove, and a presenting peptide covalently (e.g. an antigenic peptide) crowplex (I) comprising an MHC molecule, where (I) is capable of modulating the cartivity of a T call. Also included are a DNR construct codaing for the complex, where the MHC molecule is a class II MHC (e.g. mouse I-Ad or I-As or Human HLA-DRI (human leukocyee antign-DRI), a multivalent MHC fusion complex complex that modulate the activity of T cells (involving introducing peptide that can modulate the activity of T cells (involving introducing peptide that can modulate the activity of T cells (involving introducing culturing the host cells under conditions suitable for expression of the MHC fusion complex, and selecting host cells that express MHC fusion complex, complex wind protein a single recombinant expression vector complex wind protein, a single recombinant complex that modulate the activity of T cells), a single recombinant complex that modulate the activity of T cells), a single recombinant complex that codes for a T cell costimulatory factor and the alpha and beta chains of the MHC fusion complex. The DNA constructs can contain beta chains of the fusion complex. Also included are inducing an immune complex of the fusion complex. Also included are inducing an immune cereponse in a mammal (including vaccinating a mammal about peptide that contains a transmembrance of the mammal a DNA sequence comprising an expression vector. Only sequence coding for a fusion complex which is a single chain fusion complex of the mammal and a DNA sequence comprising an expression of the mammal and the contains at transmembrance of the mammal and the contains at transmembrance of the mammal and the contains and is covalently linked to the MHC protein, or DNA sequence coding for the fusion complex which is a single chain fusion complex which is a single chain fusion complex which is a single chain fusion compl
  Novel major histocompatibility complex fusion complex having presenting peptide covalently linked to MHC molecule containing peptide-binding groove, used for suppressing immune response in multiple sclerosis,
   Edwards AC;
   u;
o
   T, 0
   Grammer S,
   Seguence 1385 BP; 316 A; 383 C; 399 G; 287
  Example 17; Fig 27; 126pp; English.
   Rhode PR, Weidanz JA,
   94US-00283302.
95US-00382454.
97US-00776084.
   06-JUL-2001; 2001US-00900379
  Chavaillaz P, Jiao JJJ;
  (DADE-) DADE INT INC.
  2003-341126/32.
  P-PSDB; ABU72106
  US2002198144-A1.
   29-JUL-1994;
01-FEB-1995;
17-JAN-1997;
  26-DEC-2002
   Synthetic.
  allergies.
   HC,
   Wong
Mouse MHC I-Ad/Ova 323-339 synthetic gene SSC1.
```

No.:

```
MHC; major histocompatibility complex; PCR; polymerase chain reaction; T cell activity modulator; antagonist; immune disorder; allergy; multiple sclerosis; insulin-dependent diabetes mellitus; rheumatoid arthritis; myasthenia gravis; ds.
  Major histocompatability complex fusion complex for modulating T cell activity - used in the treatment of immune disorders, e.g. multiple sclerosis, IDDM and rheumatoid arthritis.
   168. .452
/*tag= e
/label= I-Ad beta1
/*tace= "murine MHC class II I-Ad gene beta-1 domain"
453. .734
/*tag= f
        SCT1-derived single chain gene encoding MHC fusion complex
  label= I-Ad beta2
/note= "murine MHC class II I-Ad gene beta-2
735. .806
   /*tag= c
/label= OVA 323-339
/note= "chicken ovalbumin residues 323-339"
138. .167
/*tag= d
/note= "10 residue linker peptide"
   /*tag="24 residue peptide linker"
// note= "24 residue peptide linker"
// note= "24 residue peptide linker"
// label= I-Ad_alpha1
// tag= 1 1-Ad_alpha2
// tag= 1 1-Ad_alpha2
// note= "murine MHC class II I-Ad gene
1353. 1505
// tag= 1
// note= "murine MHC class II I-Ad gene
1353. 1505
// tag= 1
// note= "murine MHC class II I-Ad gene
domain"
  Grammer
   Example 17; Fig 28; 210pp; English
  Wong HC, Rhode PR, Weidanz JA,
Chavaillaz P, Jiao J;
  94US-00283302.
95US-00382454.
   95WO-US009816
  (DADE-) DADE INT INC.
   WPI; 1996-129343/13.
P-PSDB; AAR98906.
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  29-JUL-1994;
01-FEB-1995;
  misc_feature
  misc_feature
  misc_feature
  misc_feature
   misc_feature
   misc_feature
  misc_feature
   31-JUL-1995;
   WO9604314-A1
   15-FEB-1996.
   sig_peptide
  Synthetic
            /ector
  Key
140
   ValAlaileSerLeuSerArgThrGluAlaLeuAsnHisHisAsnThrLeuValCysSer 180
   ValThraspPheTyrProAlaLys1leLysValArgTrpPheArgAsnGlyGlnGluGlu 200
   220
  240
   TyrAspSerAspValGlyGluTyrArgAlaValThrGluLeuGlyArgProAspAlaGlu 120
   344
   141 ArgHisAsnTyrGluGlyProGluThrSerThrSerLeuArgArgLeuGluGlnProAsn 160
   SerLeuLysSerProlleThrValGluTrpArgAlaGlnSerGluSerAlaArgSerLys 260
  164
   165 GGGGGAAACTCCGAAAGGCATTTCGTGGTCCAGTTCAAGGGGAGGCTACTACAACAAC 224
   464
   524
   584
   644
  20
  9
   80
  21 SerSerProGlyThrGluGlyGlyAsnSerIleCysPheSerProSerLeuGluHisPro 40
  MetAlaLeuGlnIleProSerLeuLeuLeuSerAlaAlaValValLeuMetValLeu
  6 ATGGCTCTGCAGATCCCCCAGCCTCCTCTCAGCTGCTGCTGGTGGTGCTGATGGTGTTG
  41 IleValValSerGlySerTrpAspGlyGlyGlyGlyGlyGerLeuValProArgGlySerGly
   120 ATCAACGAAGCTGGTCGTGCTAGCGGAGGGGGCGGAAGC--------GGCGGA
  GlyGlyGlySerGluArgHisPheValValGlnPheLysGlyGluCysTyrTyrThrAsn
   TyrTrpAsnSerGlnProGluIleLeuGluArgThrArgAlaGluValAspThrAlaCys
   201 ThrValGlyValSerSerThrGlnLeulleArgAsnGlyAspTrpThrPheGlnValLeu
   105 AGCCTGAAGAGCCCCCATCACTGTGGAGTGG------ACTAGTGGTGGTGGTGGCAGC
  GTCGCCATCTCCCTGTCCAGGACAGAGCCCCTCAACCACCACAAAAACACTCTGGTCTGTTCG
           1385
227
4
24
10
         Length:
Matches:
Conservative:
Mismatches:
Indels:
Gaps:
  (1-1385)
  10-048-116B-6 (1-306) x ACA60742
  DNA; 1508
  Glyglyglyglyser 265
   1.3e-98
1145.00
87.2%
85.7%
   (first entry)
   AAT17587 standard;
                             :ent Similarity:
: Local Similarity:
:y Match:
   26-SEP-1996
 nment Scores
```

191 465 181

101

61

121

alpha-transmembrane

gene

Edwards

ŝ

756

AAT17587

261

241

domain

gene alpha-2

gene alpha-2

domain"

AAT17587 encodes a murine MHC fusion complex capable of modulating T cell activity encoded by the vector SCT1. The MHC fusion complex comprises at least one MHC molecule containing a peptide-binding groove and a presenting peptide covalently linked to the MHC molecule and opt. a transmembrane domain. DNA encoding a MHC fusion complex may be cloned into a host cell to express the complex. The transformed cells may then be used to identify peptides that modulate, pref. antegonies, T cell activity. DNA encoding a MHC fusion complex or a single chain fusion molecule may be used to vaccinate a mammal against a targeted disorder. The fusion complexes may be used to suppress an immune response in an animal suffering from an immune disorder e.g. multiple sclerosis, insulindependent diabetes mellitus, rheumacoid arthritis, myasthenia gravis or chronic allergies. The complexes may also be used in the treatment of livestock and pets such as casts and dogs. The MHC fusion complexes can be produced such that they contain a single antignic peptide including one of known structure, additionally a wide range of peptides can be presented for T cell interaction

Sequence 1508 BP; 337 A; 414 C; 440 G; 317 T; 0 U; 0 Other;

| 5) x AAT17587 (1-1508) | uGlnIleProSerLeuLeuLeuSerAlaAlaValValValLeuMetVal<br> | oG1yThrG1uG1yG1yAanSer1leCysPheSerProSerLeuG1uH1s<br> | 41 IleValValSerGlySerTrpAspGlyGlyGlySerLeuValProArgGlySer |
|------------------------|-------------------------------------------------------|-------------------------------------------------------|-----------------------------------------------------------|
| × AA                   | GlnIle<br>      <br> CAGATC                           | GlyThr<br>   <br>AGGACC                               | Sergly                                                    |
| )-048-116B-6 (1-306    | 1 MetalaLeu<br>        <br>6 ATGGCTCTG                | 21 SerSerPro<br>        <br>66 AGCAGCCCA              | 41 Ilevalval                                              |
|                        | )-048-116B-6 (1-306) x AAT17587 (1-1508)              | 1.048-116B-6 (1-306) x AAT17587 (1-1508)              | 1.048-116B-6 (1-306) x ANT17587 (1-1508)                  |

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TyrAspSerAspValGluGyrArgAlaValThrGluLeuGlyArgProAspAlaGlu 120
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  TyrTrpAsnSerGlnProGluIleLeuGluArgThrArgAlaGluValAspThrAlaCys 140
  404
   141 ArgHisAsnTyrGluGlyProGluThrSerThrSerLeuArgArgLeuGluGlnProAsn 160
   464
   ValalaileSerLeuSerargThrGlualaLeuAsnHisHisAsnThrLeuValCysSer 180
   524
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  584
  ---- 119
  ------GGCGGA 164
  165 gegegaaacricceaaagecarrrceregrecaerrcaagesceaerecracracaeceae 224
  9
20
                        65
   sPro 40
   80
              Iren
   rGly
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  121
   345
  405
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  181
  525
   201
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644

```
Single chain major histocompatibility complex comprising linked alpha and beta chains - useful for suppressing an immune response to an auto:immune disease, e.g. multiple sclerosis, rheumatoid arthritis, diabetes mellitus, etc.
                   704
                                     241 SerleulysSerProlleThrValGluTrpArgAlaGlnSerGluSerAlaArgSerLys 260
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  Sequence 1508 BP; 337 A; 413 C; 441 G; 317 T; 0 U; 0 Other;
   The present sequence was used in the construction of major histocompatibility complex (MHC) fusion complexes
   1508
227
4
24
10
   Length:
Matches:
Conservative:
Mismatches:
Indels:
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  HB.
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87.2%
85.7%
70.7%
   SCT1 single chain gene; ss.
  27-MAR-1998 (first entry)
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Best Local Similarity:
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  31-JAN-1996;
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  Synthetic
   756
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DB:
  RESULT 14
  Score:
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HC; Wong

, , Jiao

Burkhardt M,

Acevedo J,

Rhode PR,

97US-00960190

29-OCT-1997;

(SUNO-) SUNOL MOLECULAR CORP.

```
240
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  344
   140
  181 ValThrAspPheTyrProAlaLysIleLysValArgTrpPheArgAsnGlyGlnGluGlu 200
   220
   SerbeuLysSerProlleThrValGluTrpArgAlaGlnSerGluSerAlaArgSerLys 260
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  284
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  524
   584
   644
  705 AGCCTGAAGAGCCCCATCACTGTGGAGTGG-----ACTAGTGGTGGCGGTGGCAGC
   645 Greargerregagargaccecrearcagogagaggreracaecergecargrogagearee 704
   Major histocompatibility complex; MHC; single chain MHC; sc-MHC; Ig; peptide binding groove; immunoglobulin; T cell receptor; immune response; immune-related disorder; antigenic peptide; fusion protein; ss.
  9
SerSerProGlyThrGluGlyGlyAsnSerIleCysPheSerProSerLeuGluHisPro
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   WO9921572-A1
  06-MAY-1999
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   241
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The invention relates to new single chain major histocompatibility complex (sc-MHC) class II complexes that comprise a peptide binding groove, and a modified class II beta 2 chain or covalantly linked class in beta 2 chain or covalantly linked immunoglobulin (Ig) light chain constant (CI) region. The MHC complexes are useful for detection and analysis of peptide ligands, pathogenic T-cells, for functional, cellular and molecular assays. They can be used to identify and isolate T cell receptor and/or MHC agonists and antagonists. They can also be used to calls involved in immune-related disorders. They can also be used to craise antibodies and to screen immune cells. It is also use in a method of suppressing an immune response immune cells. It is also use in a method of suppressing and to screen immune cells. The sc-MHC complexes comprising midlied than each of in an antipole enhanced yield. These MHC complexes also can contain single antigenic peptides readily isolated from expressing cells in significant quantities. The polyspecific MHC complexes also provide a complex. The present sequence represents a DNA encoding a single chain complex. The present sequence represents a DNA encoding a single chain complex. The present sequence represents a DNA encoding a single chain complex. The present sequence represents and contain complex. The present sequence represents a DNA encoding a single chain contain complex. The present sequence represents and contain complex. The present sequence represents a DNA encoding a single chain compared the contain complex. The present sequence represents and contain complex. The present sequence represents and contain complex. The present sequence represents and contain complex. The present sequence represents a DNA encoding a single chain contain contain contain complex. The present sequence represents and contain complex complexes and contain complex complexes and contain complexes and contain complexes and contain complexes and contain complexes and contain complexes and contain 
  100
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  Single chain major histocompatibility complex class I complexes
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TyrAspSerAspValGlyGluTyrArgAlaValThrGluLeuGlyArgProAspAlaGlu 120

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TyrTrpAsnSerGlnProGluIleLeuGluArgThrArgAlaGluValAspThrAlaCys

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13-OCT-1998;

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Recombinant proteins and molecular complexes derived therefrom, analogous to molecules involved in immune responses
Patent: WO 0.109194-A 1.08-FEB-2001;
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   541 ATCCCTTCTGATGATGACATTTATGACTGCAAGGTGGAGCACTGGGGCCTGGAGGAGCCG
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  121
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Original source text: Mus musculus (strain BALB/c, sub_species
Original source text: Mus musculus (strain BALB/c, sub_species
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The protein domains are as follows: first external protein domain
(D1) at bases 93-356; second external protein domain (D2) at bases
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antigen; cell surface glycoprotein; class II gene; glycoprotein; histocompatibility antigen; integral membrane protein; major histocompatibility complex.
Mus musculus (house mouse)
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Broolst.C.O., Mathis,D.J., Kanter,M.R., Williams,V.E. II and
McDevitt,H.O.
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1. (bases 1 to 771)

Gao, M., Wang, H. and Wang, Q.

Establishment of pIRES- I-A(d) alpha beta and stable expression of BALB/c mouse I-Ad alpha beta chain gene in NIH3T3 cell line
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Direct Submission
Submitted (29-OCT-2003) Lab of Transfusion Transfered virus,
Transfusion Institute Beijing, 27 TaiPing Road, Beijing 100039,
China
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Mus musculus
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   1 (bases 1 to 4713)
Webb,S.R., Wingvist.O., Karlsson,L., Jackson,M.R. and Peterson,P.A.
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  VIH-MGC Project URL: http://mgc.nci.nih.gov
Contact: MGC help desk
Email: cgapbs-r@mail.nih.gov
Tisaue Procurement: Jeffrey Green M.D.
Tisaue Procurement: Jeffrey Green M.D.
CDNA Library Preparation: Life Technologies, Inc.
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DNA Sequencing Dy: Baylor College of Medicine Human Genome
Sequencing Center:
Center code: BGM-HGSC
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Contact: amg@bcm.tmc.edu
Gunaratne, P.H., Garcia, A.M., Lu, X., Hulyk, S.W., Loulseged, H.,
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A.N., Gibbs, R.A.
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Mammallan Gene Collection Program Team

Generation and initial analysis of more than 15,000 full-length

human and mouse cDNA sequences

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Clark, B.R., Sharma, S.D. and Lerch, B.L.
MHC conjugates useful in ameliorating
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Clark, B.R., Sharma, S.D. and Lerch, B.L.

MHC conjugates useful in ameliorating autoimmunity

Patent: US 6451314-A 4 17-SEP-2002;

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CDNA Library Preparation: Life Technologies, Inc.
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DNA Sequencing by: Baylor College of Medicine Human Genome
Sequencing Center
Center code: BCM-HGSC
Web site: http://www.hgsc.bcm.tmc.edu/cdna/
Contact: amg@bcm.tmc.edu
Gontact: amg@bcm.tmc.edu
Kowis, C.R., Sneed, A.J., Martin, R.G., Muzny, D.M., Nanavati,
A.N., Gibbs, R.A.
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9
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  546 ATCCCTTCTGACGATGACATTTATGACTGCAAGGTGGAACACTGGGGCCTGGAGGAGCCG 605
   Mus musculus
Eukaryota, Metazoa, Chordata, Craniata, Vertebrata, Euteleostomi,
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  Direct Submission
Submitted (19-DEC-2001) National Institutes of Health, Mammalian
Gene Collection (MCC), Cancer Genomics Office, National Cancer
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  Contact: MGC help desk
Email: cgapbe.r@mail.nh.gov
Tissue procurement: Lothar Hennighausen Ph.D., Robin Humphreys
   Proc. Natl. Acad. Sci. U.S.A. 99 (26), 16899-16903 (2002) 12477932
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Email: cgapbs-r@mail.nih.gov
Tissue Procurement: Lothar Hennighausen Ph.D., Robin Humphreys
Tissue Procurement: Lothar Hennighausen Ph.D., Robin Humphreys
CDNA Library Preparation: Life Technologies, Inc.
CDNA Library Arrayed by: The I.M.A.G.B. Consortium (LLNL)
DNA Sequencing Dy: Baylor College of Medicine Human Genome
Sequencing Center
Center code: BCM-HGSC
Web site: http://www.hgsc.bcm.tmc.edu/cdna/
Contact: amg&bcm.tmc.edu
Gunaratne, P.H., Garcia, A.M., Lu, X., Hulyk, S.W., Loulseged, H.,
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   140
   366 AAGTCCCCTGTGCTGCTGGGTCAGCCCAACACCCTCATCTGCTTTGTGGACAACATCTTC 425
  485
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  181 IleProSerAspAspIspIleTyrAspCysLysValGluHisTrpGlyLeuGluGluBro 200
  141 ProProValileAsnileThrTrpLeuArgAsnSerLysSerValThrAspGlyValTyr 160
  546 Arcccrrcradacarcacarrandacracaacarcanacarcacacacacacacacaca 605
  Mus musculus
Bukaryota, Metazoa; Chordata; Craniata; Vertebrata; Buteleosťomi;
Mammalia; Butheria; Buarchoncoglires; Glires; Rodentia;
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Proc. Natl. Acad. Sci. U.S.A. 99 (26), 16899-16903 (2002)
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JOURNAL
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  PheGluProGlnGlyGlyLeuGlnAsnIleAlaAlaGluLysHisAsnLeuGlyIleLeu
  264 Trrigaeccccaaegregacrecaaaacaraecracaegaaaacacaacrregaaarcrre
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  M21931.1 GI:199493
M21931.1 GI:199493
cell surface glycoprotein; class II gene; integral membr
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Mus musculus
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gene (k haplotype),
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203
14
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Conservative:
Mismatches:
Indels:
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DB:
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LOCUS
DEFINITION
ACCESSION
VERSION
KEYWORDS
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90
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ORGANISM
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   a
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  g
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   셤
   ò
   g
   ઠે
  셤
  ò
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Mus musculus (house mouse)
Mus musculus
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Proc. Natl. Acad. Sci. U.S.A. 80 (2), 534-538 (1983)
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Submitted (05-MAY-1983)
Data kindly reviewed (05
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Patent: US 5468481-A 4 21-NOV-1995;
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Unclassified.

E 1 (bases 1 to 1508)

(S Rhode, P. R., Jalao, J. -A., Burkhardt, M. and Wong, H.C. Single chain MHC complexes and uses thereof AL Patent: US 5869270-A 122 09-FEB-1999;

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Recombinant proteins and molecular complexes derived therefrom, analogous to molecules involved in immune responses
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Kappler, J. W. and Marrack, P.

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1 (bases 1 to 1013)
Rappler, J. W. and Marrack, P.
Product and process for T cell regulation
Patent: US 5820866-A 38 13-OCT-1998;
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1 (bases 1 to 4724)

1 (bases 1 to 4724)

Webb,S.R., Winqvist,O., Karlsson,L., Jackson,M.R. and Peterson,P.A.

WHC class II antigen-presenting systems and methods for activating CD4+ T cells

CD4+ T cells

Patent: US 6355479-A 8 12-MAR-2002;

Location/Qualifiers

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1 (bases 1 to 1382)
Rhode, P.R., 'Jako, J.-A., Burkhardt, M. 'Amt molecules and uses thereof
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1 (bases 1 to 1382)
Rhode,P.R., Jiao,J.-A., Burkhardt,M. and Wong,H.C. Single chain MMC complexes and uses thereof Patent: US 5869270-A 123 09-FEB-1999;
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Inde, P.R., Jiao, J.A., Burkhardt, M. and M. Complexes and uses thereof Patent: BP 1526141-A 123 27-APR-2005;
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  1 (bases 1 to 1385)
Rhode, P.R., Jiao, J.A., Burkhardt, M. and Wong, H.C.
Single chain MHC complexes and uses thereof
Patent: US 5869270-A 121 09-FEB-1999;
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   Chavaillaz, P.A., Edwards, A.C., Grammer, S., Jiao, J.A., Rhode, P.R., Weidanz, J.A. and Wong, H.C.
Weidanz, J.A. and uses thereof
Patent: EP 0991477-A 133 03-MAY-2000;
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  Rhode, P.R., Jiao, J.A., Burkhardt, M. and Wong, H.C. MHC complexes and uses thereof
Patent: BP 1526141-A 121 27-APR-2005;
Altor BioScience Corporation (US)
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Aat04262 Hybrid IA
Adg31225 I-Ab(beta
  June 30, 2006, 01:23:04; Search time 522.277 Seconds (without alignments) 6007.369 Million cell updates/sec
   Pred. No. is the number of results predicted by chance to have a score greater than or equal to the score of the result being printed, and is derived by analysis of the total score distribution.
  US-10-048-116B-6_COPY_1_300
1572
1 MALQIPSLLLSAAVVVLMVL......KKKNAQLKWKLQALKKKLAQ 300
   Description
               GenCore version 5.1.9
Copyright (c) 1993 - 2006 Biocceleration Ltd.
  - nucleic search, using frame_plus_p2n model
  Total number of hits satisfying chosen parameters:
  5244920 seqs, 3486124231 residues
  SUMMARIES
   Post-processing: Minimum Match 0%
Maximum Match 100%
Listing first 45 summaries
  AAF55099
AAT04262
ADQ31225
  Xgapop 10.0 , Xgapext
Ygapop 10.0 , Ygapext
Fgapop 6.0 , Fgapext
Delop 6.0 , Delext
  geneseqn2006s:*
   유
  Minimum DB seq length: 0
Maximum DB seq length: 200000000
  Query
Match Length DB
  921
893
945
   BLOSUM62
   Command line parameters:
  100.0
  1122:::::
  1572
1255.5
1222
  Perfect score:
   Scoring table:
   Score
```

Database :

٠ و Result

220 9 240 260

280 840

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661 ercarecrecacareaccecrearcaceacacacereracacerecearerecarecearece 720
  AGCCTGAAGAGCCCCATCACTGTGGAGTGGAGGCACAGTCCGAGTCTGCCCGGAGCAAG 780
   LyslyslysAsnAlaGlnLeuLysTrpLysLeuGlnAlaLeuLysLysLysLeuAlaGln 300
   ValAlaIleSerLeuSerArgThrGluAlaLeuAsnHisHisAsnThrLeuValCysSer
                         ValThraspPheTyrProAlaLysIleLysValArgTrpPheArgAsnGlyGluGlu
   GTGACAGATTTCTACCCCAGCCAAGATCAAAGTGCGCTGGTTCAGGAATGGCCAGGAGGAG
   ThrvalGlyvalSerSerThrGlnLeuIleArgAsnGlyAspTrpThrPheGlnValLeu
   ValMetLeuGluMetThrProHisGlnGlyGluValTyrThrCysHisValGluHisPro
  SerLeulysSerProlleThrValGluTrpArgAlaGlnSerGluSerAlaArgSerLys
  GlyGlyGlyGlySerThrThrAlaProSerAlaGlnLeuLysLysLysLeuGlnAlaLeu
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  primer #261 (see AAT04260)"
  /*tag= a
/note= "probable primer binding site (primer #233)"
complement(45. .74)
  /*tag= h
/note= "probable primer binding site (primer #270)"
511. .825
   /*tag= g
/note= "primer #333 (see AAT04263) binding site"
/71. .389
   Polymerase chain reaction; PCR; primer; amplify; major histocompatability complex; MHC; T-cell receptor; TCR; autoimmune disease; immunodeficiency disease; immune response; immunoproliferation disease; graft-host rejection; therapy; 88.
  product= "hybrid IA beta chain"
   /*tag= b
/note= "binding site for
   /*tag= d
/note= "leader region"
complement(119. .172)
   Location/Qualifiers
   complement (199. /*tag= g
  BP.
  AAT04262 standard; DNA; 893
   U
  16-APR-1996 (first entry)
   Hybrid IA beta chain gene
   61. .828
  61. .141
   511. .o.
/*tag=
   /*tag=
  misc_feature
   primer_bind
   primer_bind
   primer_bind
  primer_bind
   sig_peptide
   primer_bind
   mat_peptide
  Synthetic.
  481
  721
  841
   201
  601
  281
  161
   181
  541
   221
  241
   261
  AAT04262;
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  The specification describes soluble recombinant proteins that comprise at least a dimer formed from the alpha and beta-chains of MHC (major histories) and comparable to complex) Class I and II molecules in which at least cone chain has, attached to its C-terminus, at least part of the Fc region of an immunoglobulin. The recombinant proteins, when linked to an antigenic peptide, are used to count and/or purify antigen-reactive T lymphocytes and to characterize their phenotype, e.g. in preclinical evaluation of vaccines. They are also used as immunostimulants, particularly for vaccine development (against infections and tumours), to count and determine phenotype of autoreactive T cells in subjects with, or at risk of developing, autoimmune diseases, e.g. for staging or evaluating treatments, and (to purify and/or enrich Ag-reactive T cells from cell cultures or patient samples, for use in subsequent curative or preventative cellular therapy. The present sequence encodes a recombinant protein of the invention, comprising a beta chain of MHC molecules
   240
   100
  120
  360
   140
  420
   480
   120
  180
   300
  ArgHisAsnTyrGluGlyProGluThrSerThrSerLeuArgArgLeuGluGlnProAsn 160
   9
   9
  40
  80
   ATGCTCTGCAGATCCCCAGCCTCCTCTCAGCTGCTGTGGTGGTGGTGATGGTGGTGTTG
  SerSerProGlyThrGluGlyGlyAsnSerIleCysPheSerProSerLeuGluHisPro
  AGCAGCCCCGGGACTGAGGCGGAAACTCCCATCTGCTTCTCGCCGTCGCTGGAGCACCCG
  ATCGTGGTGCCCGCCAGCTGGGACCGAGGTGGGGGCTCACTAGTGCCCCCGAGGCTCTGGA
   GlyThrGlnArglleArgLeuValThrArgTyrIleTyrAsnArgGluGluTyrValArg
  TyrAspSerAspValGlyGluTyrArgAlaValThrGluLeuGlyArgProAspAlaGlu
  301 TACGACAGCGACGTGGGCGAGTACCGCGGTGACCGAGCTGGGGCGGCCAGACGCCGAG
   TACTGGAACAGCCAGCCGGAGATCCTGGAGCGAACGCGGGCCCAAGGTGGACACGGCGTGC
  41 IleValValSerGlySerTrpAspGlyGlyGlyGlySerLeuValProArgGlySerGly
  GlyGlyGlySerGluArgHisPheValValGlnPheLysGlyGluCysTyrTyrThrAsn
  GGGACGCAGCGCATACGGCTCGTGACCAGATACATCTACAACCGGGAGGAGTACGTGCGC
   TyrTrpAsnSerGlnProGluIleLeuGluArgThrArgAlaGluValAspThrAlaCys
   MetAlaLeuGlnIleProSerteuLeuLeuSerAlaAlaValValValLeuMetValLeu
  New soluble recombinant protein, useful e.g. as immunostimulant, comprises dimeric major histocompatibility complex molecule fused to
  Seguence 921 BP; 214 A; 265 C; 286 G; 156 T; 0 U; 0 Other;
  921
300
0
0
  US-10-048-1168-6_COPY_1_300 (1-300) x AAF55099 (1-921)
   Length:
Matches:
Conservative:
Mismatches:
Indels:
  Example 1; Page 34-35; 43pp; French.
(CNRS ) CNRS CENT NAT RECH SCI
   2.21e-142
1572.00
100.0%
100.0%
  Malherbe
   immunoglobulin Fc region.
  WPI; 2001-182944/18.
  Percent Similarity:
Best Local Similarity:
   P-PSDB; AAB67481
  Glaichenhaus N,
   Alignment Scores:
   241
  21
   19
   121
   181
  361
  61
   81
  101
   121
  141
  421
  Query Match:
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285
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  345
   120
  405
   140
  465
   160
  525
   180
  585
   200
  645
  705
   240
  765
   80
   I-Ab(beta)-Cholera toxin B subunit-leucine zipper (LZ)-BirA fusion cDNA
   CTB; 88;
   TyrAspSerAspValGlyGluTyrArgAlaValThrGluLeuGlyArgProAspAlaGlu
   GlyGlyGlySerGluArgHisPheValValGlnPheLysGlyGluCysTyrTyrThrAsn
                             226 GGTGGAGGTCCGAAAGGCATTTCGTGGTCCAGTTCAAGGGCGAGTGCTACTACACCCAAC
   GlyThrGlnArglleArgLeuValThrArgTyrIleTyrAsnArgGluGluTyrValArg
   141 ArgHisAsnTyrGluGlyProGluThrSerThrSerLeuArgArgLeuGluGlnProAsn
   AGACACAACTACGAGGGCCGGAGACCAGCACCTCCCTGCGGGGGTTGAACAGCCCAAT
   GTCGCCATCTCCCTGTCCAGGACAGAGCCCCTCAACCACCACAAAAAACTCTGGTCTGTTCG
   ThrValGlyValSerSerThrGlnLeu1leArgAsnGlyAspTrpThrPheGlnValLeu
   ValMetLeuGluMetThrProHisGlnGlyGluValTyrThrCysHisValGluHisPro
   286 GGGACGCAGCGCATACGGCTCGTGACCAGATACATCTACAACCGGGAGGAGTACGTGCGC
  TyrTrpAsnSerGlnProGluIleLeuGluArgThrArgAlaGluValAspThrAlaCys
  TACTGGAACAGCCAGCCGGAGATCCTGGAGCGAACGCGGGGCCGAGGTGGACACGGCGTGC
   ValAlaIleSerLeuSerArgThrGluAlaLeuAsnHisHisAsnThrLeuValCysSer
   ValThrAspPheTyrProAlaLysIleLysValArgTrpPheArgAsnGlyGluGlu
   GTGACAGATTTCTACCCAGCCAAGATCAAAGTGCGCTGGTTCAGGAATGGCCAGGAGGAG
   ACAGTGGGGGTCTCATCCACACACTTATTAGGAATGGGGACTGGACTTCCAGGTCCTG
   GTCATGCTGGAGATGACCCCTCATCAGGAGAGCTCTACACCCTGCCATGTGGAGCATCCC
   /*tag= a
/product= "I-Ab(beta)-Cholera toxin B subunit (CTB)-
leucine zipper (LZ)-BirA fusion cDNA"
  class II major histocompatibility complex; MHC; CD4+ T-cell det.
flow cytometry; mucous membrane invasive antigen;
I-Ab(beta)-cholera toxin B subunit-leucine zipper-BirA fusion;
   (SENT-) SENTAN KAGAKU GIJUTSU INCUBATION CENT
  Location/Qualifiers
  BP.
  ADQ31225 standard; cDNA; 945
  03-DEC-2002; 2002JP-00351818.
   03-DEC-2003; 2003JP-00404367
  (first entry)
  Vibrio cholerae.
Unidentified.
   JP2004196789-A.
  07-0CT-2004
   101
   61
  406
  466
   191
   181
  646
   AD031225;
   81
   121
  526
   201
   221
  904
   241
  994
   ADQ3122
   RESULT
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  This sequence represents a hybrid IA beta chain gene, containing the chicken ovalbumin peptide (cOVA). This sequence was used in the construction of a hybrid IA alpha beta dimer. The encoded protein (pIAd-OVA) was found to be more stable than the IA alpha beta dimer. The stability was decreased by the addition of a MHC groove specific binding peptide (e.g. see AAR82527, AAR82528 and AAR82511), compared to an increase seen on the addition of a MHC binding peptide to IE k/dMCC. These complexes may be used to regulate an immune response. The complexes are capable of being recognised by a TCR alone or in combination with additional MHC proteins. These complexes are useful for therapeutic additional MHC proteins. These complexes are useful for therapeutic purposes and experimental purposes. They can also be used as reagents the treatment of diseases including autoimmune diseases, immunodeficiency diseases, immunoproliferation diseases, and graft-host rejection
  168
   225
   Peptide-MHC complex comprising antigenic peptide, linker and MHC segment - useful as reagents for the treatment of diseases including auto-immune diseases, immuno-stimulatory diseases or graft-host rejection.
  40
  9
  ---GAGATCAATGAGGCTGGCAGAGGAGGTGGGGGCTCACTAGTGCCCCGAGGCTCTGGA
  SerSerProGlyThrGluGlyGlyAsnSerIleCysPheSerProSerLeuGluHisPro
  41 IleValValSerGlySerTrpAspGlyGlyGlyGlySerLeuValProArgGlySerGly
   MethlaLeuGlnIleProSerLeuLeuLeuSerAlaAlaValValLeuMetValLeu
  /noce= "probable primer binding site (primer #271)"
532. .554
/*tag= k
hote= "probable primer binding site (primer #272)"
808. .836
/*tag= 1
/*tag= 1
/*tag= 1
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/*tag= 1
  'ttag= m
'note= "probable primer binding site (primer #232)"
   Sequence 893 BP; 204 A; 239 C; 275 G; 175 T; 0 U; 0 Other;
"IA beta chain beta 2 region"
  US-10-048-116B-6_COPY_1_300 (1-300) x AAT04262 (1-893)
  (NAJE-) NAT JEWISH CENT IMMUNOLOGY & RESPIRATORY.
   Length:
Matches:
Conservative:
Mismatches:
   Indels:
   Example 1; Page 53; 94pp; English
   9.23e-112
1255.50
95.4%
93.1%
  95WO-US002689
   94US-00207481
/product=
521. .550
                                      *tag=
   Marrack P;
  WPI; 1995-320543/41.
  Percent Similarity:
Best Local Similarity:
  P-PSDB; AAR82533.
  WO9523814-A1
  03-MAR-1995;
   04-MAR-1994;
  Alignment Scores:
   Kappler JW,
                 primer_bind
  primer_bind
   08-SEP-1995
   primer_bind
  primer_bind
   22
   Query Match:
DB:
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661 CTGGTCATGCTGGAGATGACCCCTCGGGGGGGAGGGCTATACACCTGTCACGTGGAGCAT 720
  601 GAGACGGTGGGGGTCTCATCCACACAGCTTATTAGGAATGGGGACTGGACCTTCCAGGTC 660
   ProSerLeuLysSerProlleThrValGluTrpArgAlaGlnSerGluSerAlaArgSer 259
   LeulyslyslysksnalaginleulysTrplysleuginalaleulyslyslysleuala 299
  Peptide-Class II major histocompatibility complex (MHC) composite, useful for detecting antigen specific CD4+ T-cell, comprises antigen peptide containing epitope of mucous membrane invasive protein, and extracellular
   /*tag= a
/product= "I-Ab(beta)-Escherichia coli heat-labile toxin
B subunit (LTB)-leucine zipper (LZ)-BirA fusion protein"
  class II major histocompatibility complex; MHC; CD4+ T-cell detection; flow cytometry; mucous membrane invasive antigen; I-Ab(beta)-heat-labile toxin B subunit-leucine zipper-BirA fusion; LTB;
                          SerValThrAspPheTyrProAlaLysIleLysValArgTrpPheArgAsnGlyGlnGlu
   541 TCAGTGACAGATTTCTACCCAGCCAAGATCAAAGTGCGCTGGTTCCGGAATGGCCAGGAG
  CTGGTTCCGCGCGGATCCACTACAGCTCCATCAGCTCAGTTGAAAAAGAAACTGCAGGCA
   LeuValMetLeuGluMetThrProHisGlnGlyGluValTyrThrCysHisValGluHis
  721 CCCAGCCTGAAGAGCCCCCATCACTGTGGAGTGGAGGGCACAGTCGTCAGCA-----GAC
   260 LysGlyGlyGlySerThrThrAlaProSerAlaGlnLeuLysLysLeuGlnAla
  I-Ab(beta)-E. coli heat-labile toxin B subunit-LZ-BirA fusion cDNA.
   (SENT-) SENTAN KAGAKU GLJUTSU INCUBATION CENT KK
   Example 3; SEQ ID NO 13; 30pp; Japanese
  Location/Qualifiers
   BP.
   ADQ31228 standard; cDNA; 915
   03-DEC-2003; 2003JP-00404367
   03-DEC-2002; 2002JP-00351818
  (first entry)
   WPI; 2004-546819/53.
P-PSDB; ADQ31227.
  coli.
   300 Gln 300
  897
  JP2004196789-A.
  五元
  Escherichia c
Unidentified.
  07-OCT-2004
  15-JUL-2004.
  region of
   240
   280
  200
   220
   775
   AD031228;
   ss; gene.
  RESULT 4
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  The invention relates to a novel class II major histocompatibility complex (MHC) antigenic peptide composite comprising a peptide containing the T-cell antigenic determinant of a mucous membrane invasive protein and the extracellular region of a class II MHC molecule or at least part of the extracellular region of the class II MHC molecule having an amino acid sequence comprising one or more deletions, substitutions or additions. The molecule of the invention may be useful for detecting an antigen-specific CD4+ T-cell by flow cytometry and for presenting a microorganism-derived mucous membrane invasive protein as an antigen. The method of the invention enables efficient detection of antigen-specific activation of CD4+ T-cells in the mucous membrane. The current sequence is that of the class II major histocompatibility complex-related I-AD(slapa)-Cholera toxin B subunit (CTB)-leucine zipper (LZ)-BirA fusion
  Peptide-Class II major histocompatibility complex (MHC) composite, useful for detecting antigen specific CD4+ T-cell, comprises antigen peptide containing epitope of mucous membrane invasive protein, and extracellular
  240
  100 ArgTyrAspSerAspValGlyGluTyrArgAlaValThrGluLeuGlyArgProAspAla 119
  360
  300
  361 GAGTACTGGAACAGCCAGCCGGGATCCTGGAGCGAACGCGGGCCGAGCTGGACACGGTG 420
   CysArgHisAsnTyrGluGlyProGluThrSerThrSerLeuArgArgLeuGluGlnPro 159
   64 cricadoricicida de contraciones de contracamentas de contracamentas de contracamente de contracamentes de contracamen
   124 GCGATCGCGGCCATCAGCATGGCGAACGGAGGTGGTCG---GGTGGAGGGGGGAAGT 180
   GluTyrTrpAsnSerGlnProGluIleLeuGluArgThrArgAlaGluValAspThrAla 139
   421 TGCAGACACAACTACGAGGGGCCGGAGACCCACACCTCCCTGCGGCGCGCTTGAACAGCCC 480
  63
   23
  79
   66
  LeuSerSerProGlyThrGluGlyGlyAsnSerIleCysPheSerProSerLeuGluHis 39
   MetAlaLeuGlnIlePro---SerLeuLeuLeuSerAlaAlaValValValLeuMetVal
   4 GIGIGICICAAAGCICCCIGGAGGIICCIACAIGGCAAAGCIGACAGIGACACIGAIGGIG
   181 GGAGGTGGAGGGTCTGAAAGGCATTTCGTGTACCAGTTCATGGGCGGAGTGCTACTTCACC
  241 AACGGGACGCAGGCGCATACGATATGTGACCAGATACATCTACAACCGGGAGGAGTACGTG
  60 GlyGlyGlyGlySerGluArgHisPheValValGlnPheLysGlyGluCysTyrTyrThr
   80 AsnGlyThrGlnArglleArgLeuValThrArgTyrlleTyrAsnArgGluGluTyrVal
   Sequence 945 BP; 230 A; 256 C; 294 G; 165 T; 0 U; 0 Other;
  945
242
14
41
   US-10-048-116B-6_COPY_1_300 (1-300) x ADQ31225 (1-945)
   Length:
Matches:
Conservative:
Mismatches:
Indels:
   Example 1; SEQ ID NO 10; 30pp; Japanese
   1.74e-108
1222.00
85.0%
80.4%
  cDNA of the invention.
                     WPI; 2004-546819/53
P-PSDB; ADQ31224.
  Similarity:
   Percent Similarity:
Best Local Similari
   region of MHC.
  Alignment Scores:
  20
   140
   120
  Query Match:
DB:
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The invention relates to a novel class II major histocompatibility complex (MHC) antigenic peptide composite comprising a peptide containing the T-cell antigenic determinant of a mucous membrane invasive protein and the extracellular region of a mucous membrane invasive protein of the extracellular region of the class II MHC molecule or at least part of the extracellular region of the class II MHC molecule having an amino acid sequence comprising one or more deletions, substitutions or additions. The molecule of the invention and be useful for detecting an antigen-specific CD4+ T-cell by flow cytometry and for presenting a microorganism-derived mucous membrane invasive protein as an antigen. The method of the invention enables efficient detection of antigen-specific activation of CD4+ T-cells in the mucous membrane. The current sequence is that of the class II major histocompatibility complex-related I-AD (alpha)-Escherichia coli hear-labile toxin B subunit (ITB)-leucine
   119
   156
  -----GGTGGTAGTGAAAGGCATTTCGTGTACCAGTTCATGGGCGAGTGCTACTTCACC 210
   159
   450
  179
  510
  199
   570
   219
   630
   LeuValMetLeuGluMetThrProHisGlnGlyGluValTyrThrCysHisValGluHis 239
  GluTyrTrpAsnSerGlnProGluIleLeuGluArgThrArgAlaGluValAspThrAla 139
  19
   63
   23
  79
  66
   LeuSerSerProGlyThrGluGlyGlyAsnSerlleCysPheSerProSerLeuGluHis 39
   ProlleValValSerGlySerTrpAspGlyGlyGlyGlySerLeuValProArgGlySer
  1 MetAlaLeuGlnIlePro---SerLeuLeuLeuSerAlaAlaValValValLeuMetVal
  SerValThrAspPheTyrProAlaLyslleLysValArgTrpPheArgAsnGlyGlnGlu
  GlyGlyGlyGlySerGluArgHisPheValValGlnPheLysGlyGluCysTyrTyrThr
  ArgTyrAspSerAspValGlyGluTyrArgAlaValThrGluLeuGlyArgProAspAla
   CysArgHisAsnTyrGluGlyProGluThrSerThrSerLeuArgArgLeuGluGlnPro
   AsnValAlalleSerLeuSerArgThrGluAlaLeuAsnHisHisAsnThrLeuValCys
  TCAGTGACAGATTTCTACCCAGCCAAGATCAAAGTGCGCTGGTTCCGGAATGGCCAGGAG
  GluThrValGlyValSerSerThrGlnLeuIleArgAsnGlyAspTrpThrPheGlnVal
   GAGACGGTGGGGGTCTCATCCACACATATTAGGAATGGGGACTGGACCTTCCAGGTC
  AsnGlyThrGlnArgileArgLeuValThrArgTyrIleTyrAsnArgGluGluTyrVal
  Sequence 915 BP; 228 A; 242 C; 271 G; 174 T; 0 U; 0 Other;
   Length:
Matches:
Conservative:
Mismatches:
Indels:
   US-10-048-116B-6_COPY_1_300 (1-300) x ADQ31228 (1-915)
  zipper (LZ)-BirA fusion cDNA of the invention.
  Gaps:
   5.91e-107
1206.00
85.0%
79.7%
   Local Similarity:
   Percent Similarity:
Best Local Similarit
  Alignment Scores:
Pred. No.:
  20
   40
  211
   391
  511
  4
   157
  80
  100
  271
   120
   331
   140
  160
  451
  180
   200
  571
   220
  Query Match:
DB:
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This nucleotide sequence comprises a PCR product obtained by amplification of mouse splenocyte cDNA using primers (see AAV12065 and AAV12066) designed for the amplification of IAA beta chain full-length cDNA. IAA alpha chain cDNA (see AAV12067) has been similarly obtained.

The IAA sequences were cloned into metallothionein promoter (see AAV12067) has been similarly obtained.

The IAA sequences were cloned into metallothionein promoter (see CDNA) the complex (WHC) class II IAA heterodimers were expressed at the cell surface of transfected Drosophila Schneider 2 (ATCC CRL or 1974) cells. The invention relates to the preparation and use of synthetic antigen presenting matrices, in particular antigen presenting cells such as insect cells that have been transfected to produce MHC antigen presenting molecules with one or more accessory molecules. The matrices are used to activate naive CD4 T cells and to shift the ongoing activation state into a preferred differentiated population of Thl or Th2 cells. Applications include the treatment of autoimmune disease, e.g.
   Major histocompatibility class II antigen; MHC class II; T cell; T lymphocyte; Th1; Th2; activation; CD4+; antigen presenting cell; APC; autoimmune disease; diabetes; multiple sclerosis; autoimmune thyroiditis; systemic lupus erythromatosus; mysthemia gravis; Crohn's disease; inflammatory bowel disease; allergy; asthma; contact sensitivity; immunotherapy; therapy; IAd beta chain; mouse; ds; circular; cyclic.
  ů
   744
  804
  299
   864
631 CTGGTCATGCTGGAGATGACCCCTCGGCGGGAGAGGTCTACACCTGTCACGTGGAGCAT 690
  - useful
   CTGGTTCCGCGCGGATCCACTACAGCTCCATCAGCTCAGTTGAAAAGAAACTGCAGGCA
  LeulyslyslysAsnAlaGlnLeulysTrpLysLeuGlnAlaLeulysLysLeuAla
   ProSerLeuLysSerProlleThrValGluTrpArgAlaGlnSerGluSerAlaArgSer
   LysGlyGlyGlyGlySerThrThrAlaProSerAlaGlnLeuLysLysLysLeuGlnAla
  Synthetic antigen presenting cell for activating CD4+ T cells - use; treat autoimmune disease, e.g. diabetes, multiple sclerosis, Crohn's disease and inflammatory bowel disease, or allergy, e.g. asthma and contact sensitivity.
   Peterson PA;
   Jackson MR,
  Example 2; Page 94-96; 141pp; English.
   Karlsson L,
   BP.
   AAV12068 standard; cDNA; 4724
   97WO-US008697.
  Murine IAd beta chain cDNA
  (first entry)
  (SCRI ) SCRIPPS RES INST
   Webb SR, Winqvist O,
   WPI; 1998-041895/04.
  867
   Gln 300
  CAG
   22-MAY-1997;
  Mus musculus
  W09746256-A1
  23-MAY-1996;
  08-JUN-1998
   11-DEC-1997.
                                  240
   260
  745
  280
   300
  865
   AAV12068
                                  8
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standard;
  WO9523814-A1
   03-MAR-1995;
  16-APR-1996
   primer_bind
   primer_bind
  08-SEP-1995.
  primer_bind
  sig_peptide
   primer_bind
   primer_bind
  primer_bind
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   primer_bind
  primer_bind
   mat_peptide
  primer_bind
   primer_bind
     AAT04269 six AAT04269; XX AAT04269; XX AAT04269; XX I G-APR-199. XX Major history was autoimmune kw amutoimmune kw amut
   Synthetic.
   949 ACAGTGGGGGTCTCATCCACACACTTATTAGGAATGGGGACTGGACCTTCCAGGTCCTG 1008
  200
  220
  240
  100
  648
  120
  708
  TyrTrpAsnSerGlnProGluIleLeuGluArgThrArgAlaGluValAspThrAlaCys 140
  768
  ArgHisAsnTyrGluGlyProGluThrSerThrSerLeuArgArgLeuGluGlnProAsn 160
   537
  537
  588
  828
  ValAlalleSerLeuSerArgThrGluAlaLeuAsnHisHisAsnThrLeuValCysSer 180
  888
   20
   SerSerProGlyThrGluGlyGlyAsnSerIleCysPheSerProSerLeuGluHisPro 40
  9
  80
diabetes, multiple sclerosis, autoimmune thyroiditis, systemic lupus erythematosus, myasthenia gravis, Crohn's disease and inflammatory bowel disease, or an allergy, e.g. asthma and contact sensitivity
   1 MetAlaLeuGlnIleProSerLeuLeuLeuSerAlaAlaValValValLeuMetValLeu
  41 IleValValSerGlySerTrpAspGlyGlyGlyGlySerLeuValProArgGlySerGly
   81 GlyThrGlnArgIleArgLeuValThrArgTyrIleTyrAsnArgGluGluTyrValArg
   TyrAspSerAspValGlyGluTyrArgAlaValThrGluLeuGlyArgProAspAlaGlu
   829 ATGGCCATCTCCCTGTCCAGGACAGAGGCCCTCAACCACCACAACACCTCTGGTCTGTTCG
  ValThrAspPheTyrProAlaLysIleLysValArgTrpPheArgAsnGlyGluGlu
  ValMetLeuGluMetThrProHisGlnGlyGluValTyrThrCysHisValGluHisPro
   GlyGlyGlySerGluArgHisPheValValGlnPheLysGlyGluCysTyrThrAsn
  538 ------TCCGAAAGGCATTTCGTGGTCCAGTTCAAGGGCGAGTGCTACTACAAC
   109 TACTGGAACAGCCGGCGAGATCCTGGAGCGAACGCGGGCCGAGGTGGACACGGCGTGC
  889 GTGACAGATTTCTACCCAGCCAAGATCAAAGTGCGCTGGTTCAGGAATGGCCAGGAGGAG
   AGACACAACTACGAGGGGCCGGAGACCAGCACCTCCCTGCGGCGGCTTGAACAGCCCAAT
  ThrValGlyValSerSerThrGlnLeuIleArgAsnGlyAspTrpThrPheGlnValLeu
   Sequence 4724 BP; 1196 A; 1194 C; 1200 G; 1134 T; 0 U; 0 Other;
  4724
228
   US-10-048-116B-6_COPY_1_300 (1-300) x AAV12068 (1-4724)
  Length:
Matches:
Conservative:
Mismatches:
Indels:
Gaps:
   511 AGCAGCCCAGGGACTGAGGGCGGAAAC-----
   1129 ATGTTGAGCĠĠĊATCĠĠĠĠĠĊ 1149
   ------GlyGlyGlyGly 264
  1.11e-101
1161.50
85.8%
85.4%
73.9%
   Percent Similarity:
Best Local Similarity:
Query Match:
DB:
   Alignment Scores:
  649
  141
   451
   21
  19
  121
  221
   261
   537
  101
  191
  181
  201
   9
   RESULT
   유
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```
(see AAT04271)"
  (see AAT04270)"
  cell;
   385. 403
/*tag= h
/*tag= h
/*tag= probable primer binding site (primer #270)"
531. 959
   #272) "
   #259) "
   (primer #271)"
   (primer #366)"
   /*tag= a
/note= "probable primer binding site (primer #76)"
complement(40. .74)
   /note= "probable primer binding site (primer #366) 1000. .1013 /*tage // hote= "probable primer binding site (primer #59)"
   /*tag= g
/note= "primer #365 (see AAT04272) binding site"
85. .403
  Major histocompatability complex; MHC; T-cell receptor; TCR; autoimmune disease; immunodéficiency disease; immune response; immunoproliferation disease; graft-host rejection; therapy; B M12.C3; pM12-IAb-Ea; 88.
   (primer
   (primer
  *tag= i
'product= "IA beta chain beta 2 region"
535. .564
   /*tag= e
/note= "binding site for primer #363
complement(177. 226)
/*tag= f
   /*tag= b
/note= "binding site for primer #362
  /*tag= j
/note= "probable primer binding site
544. .568
  note= "probable primer binding site 42. .976
   note= "probable primer binding site
  'note= "primer #364 binding site"
   (NAJE-) NAT JEWISH CENT IMMUNOLOGY & RESPIRATORY
   /*tag= c
/product= "hybrid IA beta chain"
  63. .143
/*tag= d
/note= "leader region"
complement(140. .191)
   complement (212. .266)
   Location/Qualifiers
BP.
 DNA; 1013
  95WO-US002689
   Hybrid IA beta chain gene.
  .850
  63. .959
  *tag=
  Kappler JW, Marrack P;
   (first
  WPI; 1995-320543/41.
```

235 764 255

824

```
6. 1382
/*tag= a
6. 86 6. 86
/*tag= b
/abbl= I-Ad beta chain leader
/note= "murine MHC class II I-Ad gene beta chain leader
   MHC; major histocompatibility complex; PCR; polymerase chain reaction; I cell activity modulator; antagonist; immune disorder; allergy; multiple sclerosis; insulin-dependent diabetes mellitus; rheumatoid arthritis; myasthenia gravis; ds.
CACGTGGAGCATCCCAGCCTGAAGAGCCCCCATCACTGTGGAGTGGAGGGCCACAGTCTGAG
  HisValGluHisProSerLeuLysSerProlleThrValGluTrpArgAlaGlnSerGlu
  ThrPheGlnValLeuValMetLeuGluMetThrProHisGlnGlyGluValTyrThrCys
  /label= I-Ad_alpha1
/note= "murine MHC class II I-Ad gene alpha-2 domain"
1068. .1352
   /note= "murine MHC class II I-Ad gene alpha-2 domain"
1353. .1379
   Vector SCE1-derived single chain gene encoding MHC fusion complex.
  gene beta-1
   beta-2
  /note= "chicken ovalbumin residues 323-339"
138 .167
   gene
  /*tag= d
/note= "10 residue linker peptide"
  *tag= g
note= "24 residue peptide linker"
   SerAlaArgSerLys-----GlyGlyGlyGly 264
   825 rérecerdadecaacarerrancececarcecece 860
  II I-Ad
   II I-Ad
  /*tag= e
/label= I-Ad_betal
/note= "murine MHC class
453. .734
  /note= "murine MHC class
  /*tag= c
/label= 0vA_323-339
  Location/Qualifiers
  'label = I-Ad_alpha2
  /label= I-Ad beta2
  AAT17588 standard; DNA; 1382 BP
  /note= "EE tag"
   24
.1067
/*tag= h
'labe'
   (first entry)
  sequence"
87. .137
   .452
   *tag≃
   *tag=
   *tag=
   168.
   misc_feature,
   misc_feature
   misc_feature
   misc_feature
  misc_feature
   misc_feature
   misc_feature
   WO9604314-A1
   26-SEP-1996
  misc_feature
   sig peptide
   15-FEB-1996
   Synthetic
                  645
   236
   216
   705
   AAT17588;
   765
  256
   Key
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   This sequence represents a hybrid IA beta chain gene. This sequence contains a fragment of the IE alpha chain (residues 56-73), as well as a linker and cleavage site. This sequence was transfected into a B call line (M12.C3) using plasmid pM12-IAb-Ea. It was found that the encoded sequence was expressed in these cells. Complexes such as this may be used to regulate an immune response. The complexes are capable of being recognised by a TCR alone or in combination with additional MHC proteins. These complexes are useful for therapeutic purposes and experimental purposes. They can also be used as reagents for the treatment of diseases inmunoproliferation diseases, immunodeficiency diseases,
  115
   ATGGCTCTGCAGATCCCCAGCCTCCTCCTCGGCTGCTGTGGTGGTGCTCATGGTGCTG 122
  AGCAGCCCAGGGACTGAGGGCGGAGACTCC-----GAAGCTAGCTTTGAGGCTCAG 173
   227
  ---GGGGGAAGTGGAGGTCTGAAAGGCATTTCGTGTACCAGTTCATGGGGGAG 284
   404
  ArgProAspAlaGluTyrTrpAsnSerGlnProGluIleLeuGluArgThrArgAlaGlu 135
   464
   155
   524
  175
  584
  195
   644
  196 AsnGlyGlnGluGluThrValGlyValSerSerThrGlnLeuIleArgAsnGlyAspTrp 215
                          Peptide-MHC complex comprising antigenic peptide, linker and MHC segment - useful as reagents for the treatment of diseases including auto-immune diseases, immuno-stimulatory diseases or graft-host rejection.
  40
  22
   75
  95
   MetAlaLeuGlnIleProSerLeuLeuLeuSerAlaAlaValValValLeuMetValLeu
   SerSerProGlyThrGluGlyGlyAsnSerIleCysPheSerProSerLeuGluHisPro
  GGTGCACTGGCCAACATTGCTGTCGACAAGGCTGGAGGTGGTGGATCCGGTGGA-----
   ProArgGlySerGlyGlyGlyGlySerGluArgHisPheValValGlnPheLysGlyGlu
   GluGluTyrValArgTyrAspSerAspValGlyGluTyrArgAlaValThrGluLeuGly
   CGGCCAGACGCCGAGTACTGGAACAGCCAGCCGGAGATCCTGGAGCGAACGCGGGCCGAG
   ValAspThrAlaCysArgHisAsnTyrGluGlyProGluThrSerThrSerLeuArgArg
  GTGGACACGGTGTGCAGACACTACGAGGGCCGGAGACCCACACCTCCCTGCGCGG
   LeuGluGlnProAsnValAlaIleSerLeuSerArgThrGluAlaLeuAsnHisHisAsn
   CTTGAACAGCCCAATGTCGTCATCTCCCTGTCCAGGACAGGCCCTCAACCACCACAAC
   ThrLeuValCysSerValThrAspPheTyrProAlaLysIleLysValArgTrpPheArg
   ACTCTGGTCTGCTCAGTGACAGATTTCTACCCAGCCAAGATCAAAGTGCGCTGGTTCCGG
  ------ijeValyalSerGlySerTrpAspGlyGlyGlyGlySerLeuVal
   CysTyrThrAsnGlyThrGlnArgIleArgLeuValThrArgTyrIleTyrAsnArg
   Sequence 1013 BP; 220 A; 272 C; 327 G; 192 T; 0 U; 2 Other;
  1013
230
6
22
14
   US-10-048-116B-6_COPY_1_300 (1-300) x AAT04269 (1-1013)
  Conservative:
Mismatches:
Indels:
   Length:
Matches:
   Example 2; Page 65; 94pp; English.
  1.43e-101
1151.00
86.8%
84.6%
  Percent Similarity:
Best Local Similarity:
   Alignment Scores:
   63
  174
   26
   21
  123
  41
   9/
   285
   96
  345
  116
   405
   136
   465
   156
  525
  176
  Query Match:
DB:
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404
   464
   161 ValAlaIleSerLeuSerArgThrGluAlaLeuAsnHisHisAsnThrLeuValCysSer 180
  200
  584
  Single chain major histocompatibility complex comprising linked alpha and beta chains - useful for suppressing an immune response to an auto:immune disease, e.g. multiple sclerosis, rheumatoid arthritis, diabetes mellitus, etc.
   524
  585 ACAGTGGGGGTCTCATCCACAGCTTATTAGGAATGGGGACTGGACCTTCCAGGTCCTG 644
  ValMetLeuGluMetThrProHisGlnGlyGluValTyrThrCysHisValGluHisPro 240
   645 GTCATGCTGGAGATGACCCCTCATCAGGGAGGTCTACACCTGCCATGTGGAGCATCCC 704
  SerleulysSerProlleThrValGluTrpArgAlaGlnSerGluSerAlaArgSerLys 260
   121 TyrTrpAsnSerGlnProGluIleLeuGluArgThrArgAlaGluValAspThrAlaCys
                                   345 TACTGGAACAGCCAGCCGGAGATCCTGGAGCGAACGCGGGCCCGAGGTGGACACGCGCTGC
  141 ArgHisAsnTyrGluGlyProGluThrSerThrSerLeuArgArgLeuGluGlnProAsn
  465 GICGCCATCTCCCTGTCCAGGACAGAGGCCCTCAACCACCACCACAACACTCTGGTCTG
  ValThraspheTyrProalaLyslleLysValArgTrpPheArgAsnGlyGluGlu
   525 GTGACAGATTTCTACCCAGCCAAGATCAAAGTGCGCTGGTTCAGGAATGGCCAGGAGGAG
   Thr ValGly ValSerSerThr Gln Leu Ile ArgAsn Gly AspTrp Thr Phe Gln ValLeu
   major histocompatibility complex; MHC; fusion complex;
  Ë,
   Example 17; Page 140-141; 217pp; English
   Wong
   Location/Qualifiers
  Σ
  Rhode PR, Jiao J, Burkhardt
   261 GlyGlyGlyGlySer 265
   756 GGCGGTGGTGCTTCC 770
  AAT86989 standard; DNA; 1382
  97WO-US001617.
  96US-00596387
   Construction, major histoco
SCE1 single chain gene, ss.
  (first entry)
  6. .1382
/*tag= a
  SCE1 single chain gene.
   (DADE-) DADE INT INC.
  WPI; 1997-402555/37.
  P-PSDB; AAW29214
  27-MAR-1998
  30-JAN-1997;
  31-JAN-1996;
  WO9728191-A1
   07-AUG-1997.
   Synthetic.
  241
  181
   201
  221
  705
   AAT86989;
  RESULT 8
   Key
   AAT86989
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   AAT17588 encodes a murine MHC fusion complex capable of modulating T cell activity encoded by the vector SCEI. The MHC fusion complex comprises at least one MHC molecule containing a peptide-binding groove and a presenting peptide covalently linked to the MHC molecule and opt. a transmembrane domain. DNA encoding a MHC fusion complex may be cloned into a host cell to express the complex. The transformed cells may then be used to identify peptides that modulate, pref. antagonise, T cell activity. DNA encoding a MHC fusion complex or a single chain fusion complexes may be used to suppress an immune response in an animal suffering from an immune disorder e.g. multiple sclerosis, insulindependent diabetes may be used to suppress an immune response in an animal suffering from an immune disorder e.g. multiple sclerosis, insulindependent diabetes melliture, rheumatoid arthritis, myasthenia gravis or chronic allergies. The complexes may also be used in the treatment of livestock and pets such as cats and dogs. The MHC fusion complexes can be proceded such that they conteain a single antigenic peptide including one of known structure, additionally a wide range of peptides can be presented for T cell interaction
  66 AGCAGCCCAAGGACCTTAAGTATCTCTCAGGCTGTTCACGCTGCTCACGCTGAA----- 119
  120 ATCAACGAAGCTGGTCGTGCTAGCGGAGGGGGGGGAAGC--------GGCGGA 164
  165 GGGGGAAACTCCGAAAGGCATTTCGTGGTCCAGTTCAAGGGCGAGTGCTACTACACCAAC 224
  GlyThrGlnArglleArgLeuValThrArgTyrIleTyrAsnArgGluGluTyrValArg 100
   225 GGGACGCAGCGCATACGGCTCGTGACCAGATACATCTACAACCGGGAGGAGTACGTGCGC 284
  101 TyrAspSerAspValGlyGluTyrArgAlaValThrGluLeuGlyArgProAspAlaGlu 120
  9
  80
  6 Argecrérécadarecechagecrecrerendes de la Argecra de la A
   21 SerSerProGlyThrGluGlyGlyAsnSerIleCysPheSerProSerLeuGluHisPro 40
   1 MetAlaLeuGln11eProSerLeuLeuLeuSerAlaAlaValValValLeuMetValLeu
  41 IleValValSerGlySerTrpAspGlyGlyGlyGlyGlySerLeuValProArgGlySerGly
  GlyGlyGlySerGluArgHisPheValValGlnPheLysGlyGluCysTyrTyrThrAsn
  Major histocompatability complex fusion complex for modulating T celactivity - used in the treatment of immune disorders, e.g. multiple sclerosis, IDDM and rheumatoid arthritis.
  Sequence 1382 BP; 320 A; 374 C; 404 G; 284 T; 0 U; 0 Other;
  Edwards AC;
  US-10-048-116B-6_COPY_1_300 (1-300) x AAT17588 (1-1382)
  Length:
Matches:
Conservative:
Mismatches:
Indels:
  Grammer S,
  Example 17; Fig 29; 210pp; English.
   Rhode PR, Weidanz JA,
: P, Jiao J;
  8.31e-101
1145.00
87.2%
85.7%
72.8%
95WO-US009816
  94US-00283302
95US-00382454
  (DADE-) DADE INT INC.
   WPI; 1996-129343/13.
  Similarity:
   P-PSDB; AAR98907.
   Wong HC, Rhod
Chavaillaz P,
   Percent Similarity:
31-JUL-1995;
  29-JUL-1994;
  01-FEB-1995;
   Alignment Scores:
   61
   81
  Best Local S:
Query Match:
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entry)

(first

16-JUN-2003

ACA60744;

ACA60744 standard; DNA; 1382

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ACA60744
  119
  GlyThrGlnArg1leArgLeuValThrArgTyrIleTyrAsnArgGluGluTyrValArg 100
  120
  140
   160
   180
   200
   220
  240
   164
   224
   284
   404
  464
  524
  584
  644
   704
  SerLeuLysSerProlleThrValGluTrpArgAlaGlnSerGluSerAlaArgSerLys 260
   755
   20
  65
   9
  80
   SerSerProGlyThrGluGlyGlyAsnSerIleCysPheSerProSerLeuGluHisPro 40
  GlyGlyGlySerGluArgHisPheValValGlnPheLysGlyGluCysTyrThrAsn
  GGGGGAAACTCCGAAAGGCATTTCGTGGTCCAGTTCAAGGGCGAGTGCTACTACACAAC
   ------GGCGGA
   GGGACGCAGCGCATACGGCTCGTGACCAGATACATCTACAACCGGGAGGAGTACGTGCGC
   TACGACAGCGACGTGGCGCGCGCGCGGTGACCGAGCTGGGGCGCGCCGAGACGCCGAG
   1 MetAlaLeuGlnIleProSerLeuLeuLeuSerAlaAlaValValValLeuMetValLeu
   41 IleValValSerGlySerTrpAspGlyGlyGlyGlySerLeuValProArgGlySerGly
  yArgProAspAlaGlu
   TyrfrpAsnSerGinFroGluileLeuGluArgThrArgAlaGluValAspThrAlaCys
  TACTGGAACAGCCAGCCGGAGATCCTGGAGCGAACGCGGGGCCCGAGGTGGACACGGGGGC
   ArgHisAsnTyrGluGlyProGluThrSerThrSerLeuArgArgLeuGluGlnProAsn
  AGACACAACTACGAGGGGCCGGAGACCAGCACCTCCCTGCGGCGGCTTGAACAGCCCAAT
   ValAlaIleSerLeuSerArgThrGluAlaLeuAsnHisHisAsnThrLeuValCysSer
   GTGGCCATCTCCCTGTCCAGGACAGAGCCCTCAACCACCACAAAAACACTCTGGTCTGTTCG
   ValThrAspPheTyrProAlaLysIleLysValArgTrpPheArgAsnGlyGlnGluGlu
  GTGACAGATTTCTACCCAGCCAAGATCAAAGTGCGCTGGTTCAGGAATGGCCAAGGAGGAG
   ThrValGlyValSerSerThrGlnLeuIleArgAsnGlyAspTrpThrPheGlnValLeu
  Acadidescrictricacacacacacacatariascaaridescacricaaccriccasercere
   ValMetLeuGluMetThrProHisGlnGlyGluValTyrThrCysHisValGluHisPro
   GTCATGCTGGAGATGACCCCTCATCAGGGAGAGGTCTACACCTGCCATGTGGAGCATCCC
  AGCAGCCCAAGGACCTTAAGTATCTCTCAGGCTGTTCACGCTGCTCACGCTGAA-
  Sequence 1382 BP; 320 A; 373 C; 405 G; 284 T; 0 U; 0 Other;
              of major
   1382
227
4
24
10
  TyrAspSerAspValGlyGluTyrArgAlaValThrGluLeuGl
  x AAT86989 (1-1382)
              construction
                             fusion complexes
   Length:
Matches:
Conservative:
Mismatches:
Indels:
  ATCAACGAAGCTGGTCGTGCTAGCGGAGGGGGGGGAAGC-
   Gaps:
            The present sequence was used in the histocompatibility complex (MHC) fusi
  US-10-048-116B-6_COPY_1_300 (1-300)
   8.31e-101
1145.00
87.2%
85.7%
72.8%
   770
  GlyGlyGlyGlySer 265
   GCCGCTGGTTCC
  Percent Similarity:
Best Local Similarity:
   Alignment Scores:
   225
   21
  120
  61
   165
  101
   285
  121
   345
   141
   161
   181
  525
  585
   645
   705
  261
   756
  81
  405
   201
  221
  241
   Query Match:
DB:
           The
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RESULT

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The invention relates to a major histocompatibility complex (WHC) fusion complex (I) comprising an MHC molecule that contains a peptide-binding groove, and a presenting peptide covalently (e.g. an antigenic peptide) linked to the MHC molecule, where (I) is capable of modulating the complex, where the MHC molecule is a class II MHC (e.g. mouse I-Ad or I-Asio nomplex comprising two or more linked complexes, identifying a complex comprising two or more linked complexes, identifying a complex comprising vectors that each contain the fusion complex bNA, culturing the host cells under conditions suitable for expression of the MHC fusion complex, and selecting host cells that expression of the complex, and selecting host cells that expression of the complex and selecting host cells that expression of the complex mHC protein, a single recombinant expression vector comprising DNA that codes for the alpha and beta chains of the MHC fusion complex. The DNA constructs can contain beta chains of the fusion complex. Also included are inducing an immune component of the fusion complex. Also included are inducing an immune component in a mammal (including vaccinating a mammal against a targeted disorder, by administering DNA sequence comprising a mammal against a targeted disorder, by administering DNA sequence comprising a mammal against a targeted disorder, by administering DNA sequence comprising a mammal administering CC of the mammal a DNA sequence comprising a transmembrane domain, and an an analysis of the mammal and pubminished and post and p
   presenting peptide that is a T cell receptor (TCR) antagonist or partial agonist and is covalently linked to the MHC protein, or DNA sequence coding for the fusion complex which is a single chain fusion molecule. The methods are useful for identifying a peptide that can modulate the activity of T cells, inducing an immune response in a mammal (including
  Novel major histocompatibility complex fusion complex having presenting peptide covalently linked to MHC molecule containing peptide-binding groove, used for suppressing immune response in multiple sclerosis,
  MHC; major histocompatibility complex; gene therapy; fusion compley peptide-binding groove; T cell modulation; class II MHC; vaccine; autoimmune disorder; multiple sclerosis; rheumatoid arthritis; insulin-dependent diabetes mellitus; myasthenia gravis; immunogen; chronic allergy; mouse; ds; I-Ad; gene.
   Ř
   Edwards
  Mouse MHC I-Ad/Ova 323-339 synthetic gene SCE1.
  s,
  Grammer
   Example 17; Fig 29; 126pp; English
  Weidanz JA,
  06-JUL-2001; 2001US-00900379
  94US-00283302
  95US-00382454
97US-00776084
   Jiao JJJ;
   (DADE-) DADE INT INC
   WPI; 2003-341126/32.
  Rhode PR,
  US2002198144-A1.
   P-PSDB; ABU72108
   Chavaillaz P,
  29-JUL-1994;
   01-FEB-1995;
17-JAN-1997;
  26-DEC-2002
   Mus sp.
Synthetic.
  allergies.
  H
   Wong
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ВР

```
DNA; 1385
      756 GGCGGTGGTGCTTCC 770
  *tag=
  (DADE-) DADE INT INC
                                      standard;
   Rhode PR,
   Chavaillaz P,
  misc_feature
   feature
  misc_feature
  feature
   misc_feature
   26-SEP-1996
   misc_feature
   misc_feature
  misc_feature
   WO9604314-A1
  31-JUL-1995;
   29-JUL-1994;
01-FEB-1995;
  sig_peptide
   15-FEB-1996
  Synthetic
  AAT17586;
                                      AAT17586
   HC,
                     10
  misc
   misc
  Wong
   Key
                     RESULT 1
   vaccinating a mammal against a targeted disorder) and for suppressing an immune response in a mammal. The disorders include an autoimmune disorder such as multiple sclerosis, insulin-dependent disbetes mealitus, rheumatoid arthritis, myasthenia gravis or chronic allergies. The present sequence encodes a mouse MHC class II I-Ad fusion complex of the
   100
   120
  160
   119
  140
   240
   164
   224
   284
  344
   404
  180
  200
  ThrValGlyValSerSerThrGlnLeulleArgAsnGlyAspTrpThrPheGlnValLeu 220
   464
  524
   584
   644
  20
   9
   80
  21 SerSerProGlyThrGluGlyGlyAsnSerIleCysPheSerProSerLeuGluHisPro 40
  66 AGCAGCCCAAGGACCTTAAGTATCTCTCAGGCTGTTCACGCTGCTCACGCTGAA-----
  120 ATCAACGAAGCTGGTCGTGCTAGCGGAGGGGGGGAAGC------GGCGGA
  GlyGlyGerGluArgHisPheValValGlnPheLysGlyGluCysTyrThrAsn
  GlyThrGlnhrg1leArgLeuValThrArgTyr1leTyrAsnArgGluGluTyrValArg
   TyrAspSerAspValGlyGluTyrArgAlaValThrGluLeuGlyArgProAspAlaGlu
   TyrTrpAsnSerGlnProGluIleLeuGluArgThrArgAlaGluValAspThrAlaCys
  ValAlaileSerLeuSerArgThrGluAlaLeuAsnHisHisAsnThrLeuValCysSer
   ValThraspPheTyrProAlaLysIleLysValArgTrpPheArgAsnGlyGlnGluGlu
   ValMetLeuGluMetThrProHisGlnGlyGluValTyrThrCysHisValGluHisPro
  MetAlaLeuGlnIleProSerLeuLeuLeuSerAlaAlaValValValLeuMetValLeu
  ilevalvalSerGlySerTrpAspGlyGlyGlyGlySerLeuValProArgGlySerGly
  GGGACGCAGCGCATACGCCTCGTGACCAGATACATCTACAACCGGGAGGAGTACGTGCGC
  ArgHisAsnTyrGluGlyProGluThrSerThrSerLeuArgArgLeuGluGlnProAsn
   GTCGCCATCTCCCTGTCCAGGACAGAGCCCCTCAACCCACCACAAACACTCTGGTCTGTTCG
   GTGACAGATTTCTACCCCAGCCAAGATCAAAGTGCGCTGGTTCAGGAATGGCCAGGAGGAG
   U; 0 Other
   x ACA60744 (1-1382)
  Length:
Matches:
Conservative:
Mismatches:
Indels:
  405 G; 284 T; 0
  A; 373 C;
   US-10-048-116B-6_COPY_1_300 (1-300)
  8.31e-101
1145.00
87.2%
85.7%
72.8%
  GlyGlyGlyGlySer 265
   BP; 320
  Percent Similarity:
Best Local Similarity:
Query Match:
DB:
   Sequence 1382
   gnment Scores
  invention
  41
   19
   165
   81
   225
  101
  285
  121
  141
   161
  525
   201
  465
   181
   221
   645
  241
  705
   261
   Alie
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   ద
  8
  g
  ઠે
  ద
  કે
  g
   8 8
   ઠે
```

```
Location/Qualifiers
6. 1385
6. 1485= a
6. 96
7.tag= b
/*tag= b
  MHC; major histocompatibility complex; PCR; polymerase chain reaction; T cell activity modulator; antagonist; immune disorder; allergy; multiple sclerosis; insulin-dependent diabetes mellitus; rheumatoid arthritis; myasthenia gravis; ds.
  note= "murine MHC class II I-Ad gene alpha-2 domain"
1353. 1382
  domain,
   domain"
  domain"
                           Vector SSC1-derived single chain gene encoding MHC fusion complex.
   alpha-2
   beta-1
  beta-2
  AC;
  gene
   gene
  gene
  Edwards
  residues
  *tag= g
note= "24 residue peptide linker"
107. 1067
  ′*tag≕ d
note≕ "10 residue linker peptide'
[68. .452
   I-Ad
   I-Ad
   I-Ad
   Π
   ï
  ŝ
   ï
   /*tag= c
/label= OVA_323-339
/note= "chicken ovalbumin
138. 167
   /*tag= h
/label= I-Ad_alphal
/note= "murine MHC class
1068. 1352
  Grammer
   /*tag= e
/label= I-Ad_beta1
/note= "murine MHC class
453. .734
   label= I-Ad_beta2
note= "murine MHC class
35. .806
  *tag= i
|abel= I-Ad_alpha2
   tag"
   e PR, Weidanz JA,
Jiao J;
   /*tag= j
/note= "6xHis
   95WO-US009816
  94US-00283302
95US-00382454
(first entry)
   sednence"
  . .137
   WPI; 1996-129343/13.
P-PSDB; AAR98905.
```

```
ValMetLeuGluMetThrProHisGlnGlyGluValTyrThrCysHisValGluHisPro 240
  SerLeuLysSerProlleThrValGluTrpArgAlaGlnSerGluSerAlaArgSerLys 260
   645 GrchrGcrGaahraccccrchrchagaaaaacrchachccrcchararaaaacarccc 704
  Single chain major histocompatibility complex comprising linked alpha and beta chains - useful for suppressing an immune response to an auto:immune disease, e.g. multiple sclerosis, rheumatoid arthritis, diabetes mellitus, etc.
  585 ACAGTGGGGGTCTCATCCACCACATATTAGGAATGGGGACTGGACCTTCCAGGTCCTG
181 ValThrAspPheTyrProAlaLysIleLysValArgTrpPheArgAsnGlyGlnGluGlu
                      525 GTGACAGATTTCTACCCAGCCAAGATCAAAGTGCGCTGGTTCAGGAATGGCCAGGAGGAG
  ThrValGlyValSerSerThrGlnLeuIleArgAsnGlyAspTrpThrPheGlnValLeu
  Construction; major histocompatibility complex; MHC; fusion complex; SSC1 single chain gene; 88.
  Sequence 1385 BP; 316 A; 383 C; 399 G; 287 T; 0 U; 0 Other;
  The present sequence was used in the construction histocompatibility complex (MHC) fusion complexes
   Length:
Matches:
Conservative:
Mismatches:
Indels:
Gaps:
  Rhode PR, Jiao J, Burkhardt M, Wong HC;
   Example 17; Page 135-137; 217pp; English.
  Location/Qualifiers
6. .1385
/*tag= a
   BP.
   8.33e-101
1145.00
87.2%
85.7%
72.8%
  GlyGlyGlyGlyser 265
   GCCGCTGCTTCC 770
   AAT86987 standard; DNA; 1385
  97WO-US001617.
  96US-00596387.
   (first entry)
   SSC1 single chain gene
   (DADE-) DADE INT INC
   WPI; 1997-402555/37.
P-PSDB; AAW29212.
  Score:
Percent Similarity:
Best Local Similarity:
Query Match:
DB:
   27-MAR-1998
  30-JAN-1997;
  31-JAN-1996;
  WO9728191-A1
   Alignment Scores:
Pred. No.:
   07-AUG-1997.
   Synthetic.
  221
  201
  AAT86987;
  241
  261
   756
  RESULT 11
   AAT8698.
                           g
  g
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  ઠે
   g
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   셤
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  ò
   원
   AAT17586 encodes a murine MHC fusion complex capable of modulating T cell activity encoded by the vector SSC1. The MHC fusion complex comprises at least one MHC molecule containing a postide-binding groove and a presenting peptide covalently linked to the MHC molecule and opt. a transmembrane domain. DNA encoding a MHC fusion complex may be cloned into a host cell to express the complex. The transformed cells may then ce used to identify peptides that medulate, pref. antagonise, T cell may then complex or a single chain fusion molecule may be used to vaccinate a mammal against a targeted disorder. The fusion complexes may be used to suppress an immune response in an animal suffering from an immune disorder e.g. multiple sclerosis, insulince chronic allergies. The complexes may also be used in the treatment of livestock and pets such as cats and dogs. The MHC fusion complexes can be produced such that they contain a single antigenic peptide including one of known structure, additionally a wide range of peptides can be presented for T cell interaction
  140
   66 AGCAGCCAAGGACCTTAAGTATCTCTCAGGCTGTTCACGCTGCTCACGCTGAA---- 119
  224
  GlyThrGlnArglleArgLeuValThrArgTyrIleTyrAsnArgGluGluTyrValArg 100
   284
  TyrAspSerAspValGlyGluTyrArgAlaValThrGluLeuGlyArgProAspAlaGlu 120
   ValAlalleSerLeuSerArgThrGluAlaLeuAsnHisHisAsnThrLeuValCysSer 180
  524
  9
  GlyGlyGlySerGluArgHisPheValValGlnPheIysGlyGluCysTyrThrAsn 80
  SerSerProGlyThrGluGlyGlyAsnSerIleCysPheSerProSerLeuGluHisPro 40
  GGGGGAAACTCCGAAAGGCATTTCGTGGTCCAGTTCAAGGGCGAGTGCTACTACACCAAC
  TyrTrpAsnSerGlnProGluIleLeuGluArgThrArgAlaGluValAspThrAlaCys
   ArgHisAsnTyrGluGlyProGluThrSerThrSerLeuArgArgLeuGluGlnProAsn
   GTGGCCATCTCCCTGTCCAGGACAGAGGCCCTCAACCACCACAACACTCTGGTCTGTTCG
         Major histocompatability complex fusion complex for modulating T cell activity - used in the treatment of immune disorders, e.g. multiple sclerosis, IDDM and rheumatoid arthritis.
  41 IleValValSerGlySerTrpAspGlyGlyGlyGlySerLeuValProArgGlySerGly
   Sequence 1385 BP; 316 A; 384 C; 398 G; 287 T; 0 U; 0 Other;
   1385
227
4
24
10
3
   US-10-048-116B-6_COPY_1_300 (1-300) x AAT17586 (1-1385)
   Length:
Matches:
Conservative:
Mismatches:
Indels:
Gaps:
   Example 17; Fig 27; 210pp; English.
   8.33e-101
1145.00
87.2%
85.7%
72.8%
  Percent Similarity:
Best Local Similarity:
Query Match:
DB:
  Alignment Scores:
Pred. No.:
  21
  61
   165
   225
  101
   285
  121
   345
  141
   405
   161
  81
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of major

chronic allergy; mouse; ds; I-Ad; gene.

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ValThraspPheTyrProAlaLyslleLysValArgTrpPheArgAsnGlyGluGlu 200
  ThrValGlyValSerSerThrGlnLeuIleArgAanGlyAapTrpThrPheGlnValLeu 220
   ValMetLeuGluMetThrProHisGlnGlyGluValTyrThrCysHisValGluHisPro 240
  TyrTrpAsnSerGlnProGluIleLeuGluArgThrArgAlaGluValAspThrAlaCys 140
  TACTGGAACAGCCAGCCGGAGATCTTGAACGCGGGCCGAGGTGGACACGCGGCTGC 404
   ArgHisAsnTyrGluGlyProGluThrSerThrSerLeuArgArgLeuGluGlnProAsn 160
  ValAlaIleSerLeuSerArgThrGluAlaLeuAsnHisHisAsnThrLeuValCysSer 180
  ACAGIGGGGGTCTCATCCACACACTTATTAGGAATGGGGACTGGACTTCCAGGTCCTG 644
  SerLeuLysSerProlleThrValGluTrpArgAlaGlnSerGluSerAlaArgSerLys 260
   GGGGGAAACTCCGAAAGGCATTTCGTGGTCCAGTTCAAGGGCGAGTGCTACTACACCAAC 224
   GlyThrGlnArglleArgLeuValThrArgTyrIleTyrAsnArgGluGluTyrValArg 100
  GGGACGCAGCGCATACGGCTCGTGACCAGATACATCTACAACCGGGAGGAGTACGTGCGC 284
  TyrAspSerAspValGlyGluTyrArgAlaValThrGluLeuGlyArgProAspAlaGlu 120
   285 TACGACAGCGACGTGGGCGAGTACCGCGCGTGACCGAGCTGGGGCGGCCAGACGCCGAG 344
   AGACACAACTACGAGGGGCCGGAGACCAGCACCTCCCTGCGGCGCTTGAACAGCCCAAT 464
  Greacagarrreracecageceagareaagreegeregereaaargeecagagaag 584
  Greatscrissasarisaccertearessasasasactraeacerteresasasarice 704
   AGCCTGAAGAGCCCCATCACTGTGGAGTGG-----ACTAGTGGTGGCGGTGGCAGC 755
   20
   41 IleValValSerGlySerTrpAspGlyGlyGlyGlySerLeuValProArgGlySerGly 60
   Areacreracadarececadeerecreerecreacreracides es
   21 SerSerProGlyThrGluGlyGlyAsnSerIleCysPheSerProSerLeuGluHisPro 40
  GlyGlyGlyGlySerGluArgHisPheValValGlnPheLysGlyGluCysTyrThrAsn
                                1 MetalaLeuGlnIleProSerLeuLeuLeuSerAlaAlaValValValLeuMetValLeu
  WHC; major histocompatibility complex; gene therapy; fusion complex; peptide-binding groove; T cell modulation; class II WHC; waccine; autoimmune disorder; multiple sclerosis; rheumatoid arthritis; insulin-dependent diabetes mellitus; myasthenia gravis; immunogen;
US-10-048-116B-6_COPY_1_300 (1-300) x AAT86987 (1-1385)
   Mouse MHC I-Ad/Ova 323-339 synthetic gene SSC1.
  ACA60742 standard; DNA; 1385 BP.
   GlyGlyGlyGlySer 265
   770
   (first entry)
   16-JUN-2003
   585
  ACA60742;
   221
  345
  161
   181
  645
   261
   756
  61
  165
   81
  225
  101
   121
  141
   405
   525
  201
   705
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  6 6 6
   셤
  8 8
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  용
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The invention relates to a major histocomparibility computed (Fig. 1) comprising an MHC molecule that contains a peptide-binding groove, and a presenting peptide covalently (e.g. an antiganic peptide) linked to the MHC molecule, where (I is capable of modulating the crivity of a T cell. Also included are a DNA construct coding for the complex, where the MHC molecule is a class II MHC (e.g. mouse I-Ad or I-As, or human HLA-DRI (human leukocyte antigen-DRI)), a multivalent MHC (asion complex comprising two or more linked complexes, identifying a peptide that can modulate the activity of T cells (involving introducing introducing introducing the host cells cloning vectors that each contain the fusion complex. And selecting host cells that express MHC fusion complex, and selecting host cells that express MHC fusion complex that modulate the activity of T cells), a single recombinant expression vector comprising DNA that codes for the alpha and beta chains of the MHC fusion complex. The DNA constructs can contain complex that modulate the activity of T cells), a single recombinant expression of the fusion complex. Also included are inducing an immune caponse in a mammal (including vaccinating a mammal against a targeted disorder, by administering DNA sequences and koxas sequence for efficient expression of the fusion complex. Also included are inducing an immune response in a mammal a DNA sequence comprising a fusion complex, or DNA sequence conding for a fusion complex. Mich is a single chain fusion molecule. Or the mammal a DNA sequence comprising an expression vector, encoding a presenting peptide that is a T cell receptor (TCR) antagonist or partial and suppressing an immune response in a mammal discovalently inducing an immune response in a mammal against a dargeted coding for the fusion complex which is a single chain fusion molecule. The methods are useful for identifying a peptide that can modulate the cannot complex which is a single capanism and an ammal against a dargeted disorder, by administry of T cell
  such as multiple sclerosis, insulin-dependent diabetes mellitus, rheumatoid arthritis, myasthenia gravis or chronic allergies. The present sequence encodes a mouse MHC class II I-Ad fusion complex of the
   invention relates to a major histocompatibility complex (MHC) fusion
   Novel major histocompatibility complex fusion complex having presenting peptide covalently linked to MHC molecule containing peptide-binding groove, used for suppressing immune response in multiple sclerosis,
   BP; 316 A; 383 C; 399 G; 287 T; 0 U; 0 Other;
   Edwards AC;
  1385
227
   Length:
Matches:
Conservative:
   Rhode PR, Weidanz JA, Grammer S,
  Example 17; Fig 27; 126pp; English.
  94US-00283302.
95US-00382454.
97US-00776084.
   8.33e-101
1145.00
87.2%
   06-JUL-2001; 2001US-00900379
  Jiao JJJ;
   (DADE-) DADE INT INC.
   WPI; 2003-341126/32
  P-PSDB; ABU72106.
   US2002198144-A1.
   Percent Similarity:
  Chavaillaz P,
  29-JUL-1994;
01-FEB-1995;
17-JAN-1997;
  26-DEC-2002
   Mus sp.
Synthetic.
  allergies.
  invention
   Wong HC,
```

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/*tag= j
/label= I-Ad_alpha-TM
/note= "murine MHC class II I-Ad gene alpha-transmembrane
domain"
  AAT17587 encodes a murine MHC fusion complex capable of modulating T cell activity encoded by the vector SCT1. The MHC fusion complex comprises at least one MHC molecule containing a peptide-binding groove and a presenting peptide covalently linked to the MHC molecule and opt.
  /label= I-Ad_beta_chain_leader
/note= "murine MHC class II I-Ad gene beta chain leader
sequence"
87. .137
   II I-Ad gene alpha-2 domain"
   Major histocompatability complex fusion complex for modulating T celactivity - used in the treatment of immune disorders, e.g. multiple sclerosis, IDDM and rheumatoid arthritis.
    T cell activity modulator; antagonist; immune disorder; allergy; multiple sclerosis; insulin-dependent diabetes mellitus; rheumatoid arthritis; myasthenia gravis; ds.
   II I-Ad gene alpha-2
  beta-1
   beta-2
  323-339
   Edwards AC;
  gene
   gene
  /*tag= c
//label= OVA_323-339
/note= "chicken ovalbumin residues
138. .167
  *tag= g
note= "24 residue peptide linker"
107. .1067
   /*tag= d
note= "10 residue linker peptide"
168. ,452
   II I-Ad
   /note= "murine MHC class II I-Ad
   Grammer S,
   /*tag= h
/label= I-Ad_alphal
/note= "murine WHC class
1068. 1352
  /note= "murine MHC class
  class
   Location/Qualifiers
6. .1508
   /*tag= i
/label= I-Ad_alpha2
/note= "murine MHC o
  /label= I-Ad betal
  /*tag= f
/label= I-Ad beta2
  Example 17; Fig 28; 210pp; English.
  Weidanz JA,
  95WO-US009816
  94US-00283302,
95US-00382454.
   1353. .1505
   .806
   /*tag=
5. .86
  *tag=
  Wong HC, Rhode PR, We
Chavaillaz P, Jiao J;
   (DADE-) DADE INT INC.
   WPI; 1996-129343/13.
  P-PSDB; AAR98906.
  misc_feature
   misc_feature
   misc_feature
  misc_feature
   misc_feature
  misc_feature
   misc_feature
  misc_feature
  31-JUL-1995;
  29-JUL-1994;
  01-FEB-1995;
  WO9604314-A1
   sig_peptide
   15-FEB-1996
   Synthetic.
   Key
    AGCAGCCCAAGGACCTTAAGTATCTCTCAGGCTGTTCACGCTGCTCACGCTGAA----- 119
  120 ATCAACGAAGCTGGTCGTCGTAGCGGAGGGGGGGGAAGC--------GGCGGA 164
   224
   100
   284
  120
   344
   140
   404
   160
  464
  180
  200
   220
  240
   9
  80
   524
   584
   644
  SerleulysSerProlleThrValGluTrpArgAlaGlnSerGluSerAlaArgSerLys 260
  AGCCTGAAGAGCCCCATCACTGTGGAGTGG-----ACTAGTGGTGGCGGTGGCAGC 755
   SerSerProGlyThrGluGlyGlyAsnSerIleCysPheSerProSerLeuGluHisPro 40
   ATGCCTCTGCAGATCCCCAGCCTCCTCCTCTCAGCTGCTGCTGGTGGTGGTGGTGGTGGTGGTG
   MetAlaLeuGlnIleProSerLeuLeuLeuSerAlaAlaValValLeuMetValLeu
  41 IleValValSerGlySerTrpAmpGlyGlyGlyGlyGlySerLeuValProArgGlySerGly
  GlyGlyGlySerGluArgHisPheValValGlnPheLygGlyGluCysTyrTyrThrAsn
  GlyThrGlnArgileArgLeuValThrArgTyrileTyrAsnArgGluGluTyrValArg
  TyrAspSerAspValGlyGluTyrArgAlaValThrGluLeuGlyArgProAspAlaGlu
   285 TACGACAGCGACGTGGGCCGAGTACCGCGCGGTGACCGAGCTGGGGCGGCCAGACGCCGGG
   ArgHisAsnTyrGluGlyProGluThrSerThrSerLeuArgArgLeuGluGlnProAsn
   ValAlalleSerLeuSerArgThrGluAlaLeuAsnHisHisAsnThrLeuValCysSer
  ValThrAspPheTyrProAlaLysIleLysValArgTrpPheArgAsnGlyGlnGluGlu
   ThrValGlyValSerSerThrGlnLeulleArgAsnGlyAspTrpThrPheGlnValLeu
  ValMetLeuGluMetThrProHisGlnGlyGluValTyrThrCysHisValGluHisPro
  MHC; major histocompatibility complex; PCR; polymerase chain reaction;
   465 GTGGCCATCTCCCTGTCCAGGACAGAGGCCCTCAACCACCACAACACTCTGGTCTGTTCG
   SCT1-derived single chain gene encoding MHC fusion complex.
   24
10
  US-10-048-116B-6_COPY_1_300 (1-300) x ACA60742 (1-1385)
  Mismatches:
Indels:
Gaps:
  Bb.
   GlyGlyGlyGlySer 265
  AAT17587 standard; DNA; 1508
  (first entry)
 85.7%
72.8%
Local Similarity:
  26-SEP-1996
  99
   101
   141
   61
  165
   81
   225
  191
  221
   241
  181
   AAT17587;
   201
  645
   705
  261
   756
           Query Match:
DB:
  Vector
  ð
  셤
   셤
  8
  ò
  쉽
   g
  à
  ò
  셤
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  임
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   ઠે
  ò
   요
  ठे
  요
   ò
   g
  ò
   g
  ò
   임
```

domain"

domain"

```
transmembrane domain. DNA encoding a MHC fusion complex may be cloned into a host cell to express the complex. The transformed cells may then be used to identify peptides that modulate, pref. antegonise, T cell activity. DNA encoding a MHC fusion complex or a single chain fusion molecule may be used to vaccinate a mammal against a targeted disorder. The fusion complexes may be used to suppress an immune response in an animal suffering from an immune disorder. So multiple sclerosis, insulindependent diabetes mellitus, rheumatcid arthritis, myssthenia gravis or chronic allergies. The complexes may also be used in the treatment of livestock and pets such as cats and dogs. The MHC fusion complexes can be produced such that they contain a single antigenic peptide including one of known structure, additionally a wide range of peptides can be presented for T cell interaction
   240
  140
  ValThrasppheTyrProAlaLyslleLysValArgTrpPheArgAsnGlyGlnGluGlu 200
   100
   160
  180
  524
  644
   164
  224
   284
   344
   404
   704
  40
   9
  80
   20
   65
  GlyGlyGlyGluArgHisPheValValGlnPheLysGlyGluCysTyrThrAsn
   GGGGGAAACTCCGAAAGGCATTTCGTGGTCCAGTTCAAGGGCGAGTGCTACTACAACC
  ArgHisAsnTyrGluGlyProGluThrSerThrSerLeuArgArgLeuGluGlnProAsn
  GTCGCCATCTCCCTGTCCAGGACAGAGGCCCTCAACCACCACAACACTCTGGTCTGTTCG
   ThrValGlyValSerSerThrGlnLeulleArgAsnGlyAspTrpThrPheGlnValLeu
   ACAGTGGGGGTCTCATCCACACACCTTATTAGAATGGGGACTGGACCTTCCAGGTCCTG
   ValMetLeuGluMetThrProHisGlnGlyGluValTyrThrCysHisValGluHisPro
   GTCATGCTGGAGATGACCCCTCATCAGGAGAGGTCTACACCCTGCCATGTGGAGCATCCC
  SerSerProGlyThrGluGlyGlyAsnSerIleCysPheSerProSerLeuGluHisPro
  AGCAGCCCAAGGACCTTAAGTATCTCTCAGGCTGTTCACGCTGCTCACGCTGAA-----
   IleValValSerGlySerTrpAspGlyGlyGlyGlyGlySerLeuValProArgGlySerGly
   GlyThrGlnArglleArgLeuValThrArgTyrIleTyrAsnArgGluGluTyrValArg
   GGGACGCAGCGCATACGGCTCGTGACCAGATACATCTACAACCGGGAGGAGTACGTGGC
  TyrAspSerAspValGlyGluTyrArgAlaValThrGluLeuGlyArgProAspAlaGlu
   AGACACAACTACGAGGGCCGGAGACCAGCACCTCCCTGCGGCGGCTTGAACAGCCCAAT
  ValalleSerLeuSerArgThrGluAlaLeuAsnHisHisAsnThrLeuValCysSer
  Sequence 1508 BP; 337 A; 414 C; 440 G; 317 T; 0 U; 0 Other;
  US-10-048-116B-6_COPY_1_300 (1-300) x AAT17587 (1-1508)
  Length:
Matches:
Conservative:
Mismatches:
Indels:
  9.35e-101
1145.00
87.2%
85.7%
72.8%
   Similarity:
  Percent Similarity:
Best Local Similarity
Query Match:
DB:
  Alignment Scores:
   Н
  9
   21
  99
  41
  120
   165
  225
  101
   285
  121
  345
   141
   405
  465
   181
   525
  201
   221
  81
  191
   61
   ..
0
   Score:
    88888888888888888%
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   g
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  8 6
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   8 8
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  ઠે
   કે
```

```
Single chain major histocompatibility complex comprising linked alpha and beta chains - useful for suppressing an immune response to an auto:immune disease, e.g. multiple sclerosis, rheumatoid arthritis, diabetes
SerLeulysSerProlleThrValGluTrpArgAlaGlnSerGluSerAlaArgSerLys 260
                   705 AGCCTGAAGAGCCCCATCACTGTGGAGTGG-----ACTAGTGGTGGCGGTGGCAGC 755
   AGCAGCCCAAGGACCTTAAGTATCTCTCAGGCTGTTCACGCTGCTCACGCTGAA----- 119
  65
   40
  SerSerProGlyThrGluGlyGlyAsnSerIleCysPheSerProSerLeuGluHisPro
   1 MetAlaLeuGlnIleProSerLeuLeuLeuSerAlaAlaValValLeuMetValLeu
   complex;
   n; major histocompatibility complex; MHC; fusion chain gene; 88.
   BP; 337 A; 413 C; 441 G; 317 T; 0 U; 0 Other;
   of major
  1508
227
4
24
10
3
   US-10-048-116B-6_COPY_1_300 (1-300) x AAT86988 (1-1508)
  the construction
   fusion complexes
  Length:
Matches:
Conservative:
Mismatches:
Indels:
   HC;
   Example 17; Page 137-139; 217pp; English.
   Wong
  Location/Qualifiers
6. .1508
/*tag= a
  The present sequence was used in histocompatibility complex (MHC)
  ΣÌ
   BP.
  Rhode PR, Jiao J, Burkhardt
  9.35e-101
1145.00
87.2%
85.7%
72.8%
   265
   AAT86988 standard; DNA; 1508
  97WO-US001617
  96US-00596387
   (first entry)
  SCT1 single chain gene
  (DADE-) DADE INT INC
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   344
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  284
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  464
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immune response;
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The invention relates to new single chain major histocompatibility complex (sc-MHC) class II complexes that comprise a peptide binding groove, and a modified class II beta 2 chain or covalently linked immunoglobulin (Ig) light chain constant (I) region. The MHC complexes are useful for detection and analysis of peptide ligands, pathogenic T-cells, for functional, cellular and molecular assays. They can be used to identify and isolate T cell receptor and/or MHC agonists and antagonists. They can also be used to calls involved in immune-related disorders. They can also be used to craise antibodies and to screen immune cells. It is also use in a method of suppressing an immune response in mammals. The sc-MHC complexes of suppressing an immune response in mammals. The sc-MHC complexes comprising modified class II beta 2 chains and/or Ig-(I regions are soluble and provide enhanced yield. These MHC complexes also can contain single antigenic peptides readily isolated from expressing cells in significant quantities. The polyspecific MHC complexes also provide a means to detect cells expressing multiple target structures with a single complex. The present sequence represents a DNA encoding a single chain
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Search completed: June 30, 2006, 01:48:14 Job time : 528.277 secs

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Recombinant proteins and molecular complexes derived therefrom, analogous to molecules involved in immune responses
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| Unclassified.  REFERENCE 1 (bases 1 to 893) AUTHORS Kappler, J. W. and Marrack, P. TITLE Product and process for T cell regulation JOURNAL Patent: US 5820866-A 25 13-OCT-1998; FRATURES 1. 893 1. 893 //organism="unknown" //mol_type="unassigned DNA" | Query Match Best Local Similarity 94.6%; Pred. No. 2.6e-191; Matches 748; Conservative 0; Mismatches 28; Indels 15; Gaps 2;                                           | 0y 1 ATGGCTCTGCAGATCCCCAGCCTCCTCAGCTGCTGTGGTGGTGGTGGTGGTGTGTGT                                                                                                                               | 0y 61 AGCAGCCCGGGACTGAGGGCGGAAACTCCATCTGCTTCTCGCCGTCGCTGGAGCACCCG 120 | Oy 121 ATCGTGGTGTCCGCAGCTGGGACGTGGGGGCTCACTAGTGCCCCGAGGCTCTGGA 180                                                                                | QY         181 GGTGGAGGCTCCGAAAGGCATTTCGTGGTCCAGTTCAAGGGCGAGTGCTACACCAAC         240           Db         226 GGTGGAGGCTCCGAAAGGCATTTCGTGGTCCAGTTCAAGGGCGAGTGCTACAACAAC         285 | Qy         241         GGGACGCAGCGCATACGGCTCGTGACCAGATACATCTACAACCGGAGGAGTACGTGCGC 300 | QY         301 TACGACGACGTGGGCGAGTACCGCGCGGTGACCGAGCCGGCGGCCGAGCCGGGG         360 TACGACGTGGGCGAGTACCGCGGGTGACCGAGCTGGGGCGGCCAGACCCGAGCTGGGCGGCCAGACCCGAGCTGGGCGGCCGAGCTGGGCGGCCGAGCTGGGCCGGGCGGCCGGGGCGGCCGGGGGCGGCCAGACCCGAGCTGGGCCGGGCGGCCGGGGCGGCCGGGGCGGCCGGGGGG | Qy         361         TACTGGBACAGCCAGAGATCCTGGAGCGAACGCGGGCCGAGGTGGACACGCGTGC         420                                                                          | QY         421 AGACACAACTACGAGGGCCGGAGACCAGCACCTCCCTGCGGCCGCTTGAACAGCCCAAT 480                                                                                                | Oy 481 GTCGCCATCTCCCTGTCCAGGACAGAGGCCTCAACCACCACAACATCTGGTCTGTTCG 540                                                                                                            | Oy 541 GTGACAGATTTCTACCCAGCCAAGATCAAAGTGCGCTGGTTCAGGAATGGCCAGGAGGG 600      | Oy 601 ACAGTGGGGGTCTCATCACACACACTTATTAGGAATGGGGACTTCCAGGTCCTG 660                    | Qy         661         GTCATGCTGGAGATGACCCCTCATCAGGGAGGAGGTCTACACCTGCCATGTGGAGCATCCC         720             | Qy         721 AGCCTGAAGACCCCATCACTGTGAGTGGAGGGCACAGTCTGCCCGGAGCAAG 780           Db         766 AGCCTGAAGAGCCCCATCACTGTGGAGTGGAGGGCACAGTCCGAGTCTGCCCGAGGAAG 825           Qy         781 GGAGGTGGAGG 791           Db         826 TAAGCATGCG 836           RESULT 3 |
|---------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|-----------------------------------------------------------------------------------------------------------------------------------------------------------------------|----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|-----------------------------------------------------------------------|---------------------------------------------------------------------------------------------------------------------------------------------------|-------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|----------------------------------------------------------------------------------------|-----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|---------------------------------------------------------------------------------------------------------------------------------------------------------------------|-------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|-----------------------------------------------------------------------------|--------------------------------------------------------------------------------------|--------------------------------------------------------------------------------------------------------------|----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
|                                                                                                                                                                                                                                                         | dy     181 GGTGGAGGCTCCGAAAGGCATTTCGTGGTCCAGTTCAAGGGCGAGTGCTACTACACCAAC     240       Db     181 GGTGGAGGCTCCGAAAGGCATTTCGTGGTCCAGTTCAAGGGCGAGTGCTACAACAACAAC     240 | Qy         241         GGGACGCAFACGCTCGTGACCAGATACATCTACAACCGGAGGAGTACGTGCGC         300           b         241         GGGACGCAFACGGCTCGTGACCAGATACATCTACAACCGGGAGGAGTACGTGCGC         300 | Qy         301         TACGACGACGACGTGGGGGGGGGGGGGGGGGGGGGGGGG        | Qy     361 TACTGGAACAGCCGGAGATCCTGGAGCGAACGCGGGCCGAGGTGGACGCGTGC 420       Db     361 TACTGGAACAGCCGGAGATCCTGGAGCGAACGCGGGCCGAGGTGGACAGGCGTGC 420 | Qy         421 AGACACAACTACGAGGGGCGGGAGACCAGCACCTCCCTGGGCGGCTTGAACAGCCCAAT 480           Db         421 AGACACAACTACGAGGGCCGGAGACCAGCACCTCCCTGCGGCGGCTTGAACAGCCCAAT 480             | Qy 481 GTCGCCATCTCCCTGTCCAGGACAGAGGCCTCAACCACCACAACATCTGGTCTGTTCG 540                  | Qy       541 GTGACAGATTICTACCCAGCCAGATCAAAGIGCGCTGGTTCAGGAAIGGCCAGGAGGAG       600         bb       541 GTGACAGATTCTACCCAGCCAAGATCAAAGIGCGCTGGTTCAGGAAIGGCCAGGAGGAG       600                                                                                         | Qy     601 ACAGTGGGGGTCTCATCCACACACACTTATTAGGAATGGGGACTGGACCTTCCAGGTCCTG     660       b     601 ACAGTGGGGGTCTCATCCACACGTTATTAGGAATGGGGACTGGACCTTCCAGGTCTCG     660 | Qy     661     GTCATGCTGGAGATGACCCCTCATCAGGGAGAGGTCTACACCTGCCATGTGGAGCATCCC     720       Db     661     GTCATGCTGGAGATGACCCCTCATCAGGGAGAGGTCTACACCTGCCATGTGGAGCATCCC     720 | Qy       721 AGCCTGAAGAGCCCCATCACTGTGAAGTGGAGGGCACAGTCCGAGTCTGCCGGAGCAAG       780         Db       721 AGCCTGAAGAGCCCCATCACTGTGGAGGGGAGGAGGAGCAGTCCGAGTCTGCCCGGAGCAAG       780 | Qy 781 GGAGGTGGAGGATCCACTACAGCTCCATCAGCTCAGTTGAAAAAGAAATTGCAAGCACTG 840<br> | Qy 841 AAGAAAAAGAACGCTCAGCTGAAGTGGAAACTTCAAGCCCTCAAGAAACTAGGCCCTCGAGAAACTGCCCCAG 900 | Oy         901 CATCATCATCATGAT         918           Db         901 CATCATCATCATCATCATCATCATCATCATCATCATCATC | RESULT 2 AR047947 AR047947 LOCUS DCFINITION Sequence 25 from patent US 5820866. ACCESSION AR047947 KEYWORDS SOURCE UNKNOWN.                                                                                                                                          |

| AR033964 Sequence AR033964 AR033964 . Unknown Unknown Unklassi 1 (base Rhode P. Single c                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       | J. J. J. J. J. J. J. J. J. J. J. J. J. J                                                                                                                                | Query Match Best Local Similarity 89.6%; Pred. No. 1.7e-165; Matches 673; Conservative 0; Mismatches 57; Indels 21; Gaps 1; | Qy         1 ATGGCTCTGCAGATCCCCAGCCTCCTCCTCAGCTGCTGGTGGTGCTGATGGTGCTG         6 0 | 9y 61 AGCAGCCCGGGAACTGAGGCGGAAACTCCATCTGCTTCTCGCGTCGCTGGAGCACCCG 120 | Oy 121 ATCGTGGTGTCCGCAGCTGGGACGGGGGCTCACTAGTGCCCCGAGGCTCTGGA 180  Db 105 GCTGCTCACGCTGAAATCAACGAAGCTGGTCGTGCTAGCGGAGGGGGGGG | QY         181 GGTGGAGGCTCCGAAAGGCATTTCGTGGTCCAGTTCAAGGGCGAGTGCTACTACACCAAC 240           DD         165 GGGGGAAACTCCGAAAGGCATTTCGTGGTCCAGTTCAAGGGCGAGTGCTACTACACCCAAC 224 | Qy         241 GGGACGCACACATACGCTCGTGACCAGATACATCTACAACCGGGAGGAGTACGTGCGC 300           L | QY         301 TACGACGACGTGGGGGGGGGGGGGGGGGGGGGGGGGGGG                                         | Qy         361 TACTGGAACAGCCAGCCGAGATCCTGGAGCGAACGCGGGCCGAGGTGGACACGCGGCTGC 420                                                                                         | QY         421 AGACACAACTACGAGGGCCGGAGACCAGCACCTCCCTGCGGCGCGTTGAACAGCCCAAT 480           DD         405 AGACACAACTACGAGGGCCGGAGACCAGCACCTCCCTGCGGCGGTTGAACAGCCCAAT 464 | Qy         481 GTCGCCATCTCCCTGTCCAGGACAGAGCCCTCAACAACACTCTGGTCTGTTCG 540           Db         465 GTCGCCATCTCCCTGTCCAGGACGAGCCCTCAACCACAACACTCTGGTCTGTTCG 524     | Qy 541 GTGACAGATTTCTACCCAGCCAAAGTGCGCTGGTTCAGGAATGGCCAGGAGGG 600                         | QY         601 ACAGTGGGGGTCTCATCACACACTTATTAGGAACTGGACTTCCAGGTCCTG         660           Db         585 ACAGTGGGGGTCTCATCCACACAGCTTATTAGGAATGGGGACTGGACCTTCCAGGTCCTG         644 | OY 661 GTCATGCTGGAGATGACCCCTCATCAGGGAGGTCTACACCTGCCATGTGGAGCATCCC 720 |
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Pred. No. 1.7e-165;
0; Mismatches 57;
   Rhode, P.R., Jiao, J.A., Burkhardt, M. and
MHC complexes and uses thereof
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  REFERENCE
   JOURNAL
  RESULT 6
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   Score 607.8; DB 2;
Pred. No. 1.7e-165;
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Unclassified.
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|     | 465 GTCGCCATCTCCCTGTCCAGAGCCCCTCAACCACCACAACACTCTGGTCTGTTCG                                                                                                             | Qy 541 GTGACAGATTTCTACCCAGCCAAGATCAAAGTGCGCTGGTTCAGGAATGGCCAGGAGGG 600 | 601<br>585 | Oy 661 GTCATGCTGGAGATCACCCCTCATCAGGGAGAGGTCTACACCTGCCATGTGGAGCATCCC 720                                                                          | RESULT 8 AR033962 LOCUS LOCUS AR033962 AR033962 LOCUS AR033962 LOCUS AR033962 LOCUS AR033962 LOCUS AR033962 LOCUS AR033962 LOCUS AR033962.1 GI:5949567 | KEYWORDS . SOURCE Unknown. ORGANISM Unknown. Unclassified. REFERENCE 1 (bases 1 to 1385) AUTHORS Rhode, P.R., Jiao, JA., Burkhardt, M. and Wong, H.C.                                              | TITLE Single chain MHC complexes and uses thereof JOURNAL Patent: US 5869270-A 121 09-FEB-1999; FEATURES Location/Qualifiers | source 1.1383<br>/organism="unknown"<br>/mol_type="unassigned DNA"                                                                                                       | Query Match 66.2%; Score 607.8; DB 2; Length 1385; Best Local Similarity 89.6%; Pred. No. 1.7e-165; Matches 673; Conservative 0; Mismatches 57; Indels 21; Gaps 1; | Qy         1 ATGGCTGCAGATCCCCAGCCTCCTCTCACTGCTGGTGGTGGTGGTGGTGGTGGTGGTG         60 | Qy 61 AGCAGCCCGGGACTGAGGGGGGAAACTCCATCTTCTCGCGTGGAGGACCCG 120                                                                 | Qy 121 ATCGTGGTGTCCGGCAGCTGGGAGGTGGGGGCTCACTAGTGCCCCGAGGCTCTGGA 180     | Qy 181 GGTGGAGGTTCCGAAAGGCATTTCGTGGTCCAGTTCAAGGCGAGTGCTACTACACCAAC 240  | QY         241 GGGACGCAGCATACGCTCGTGACCAGATACATCTACAACCGGGAGGAGTACGTGCGC 300           DD         225 GGGACGCGCATACGGCTCGTGACCAGATACATCTACAACCGGGAGGAGTACGTGCGC 284 | Qy         301 TACGACAGCGACGTACCGCGCGGGGGACTGACCGAGCTGACCGCGGCCAGACGCCGAG         360           Db         285 TACGACGTGGGCGAGTACCGCGCGGTGACCGAGCTGGGGCGGCCAGACGCCGAG         344 |                                                                        |
|-----|-------------------------------------------------------------------------------------------------------------------------------------------------------------------------|------------------------------------------------------------------------|------------|--------------------------------------------------------------------------------------------------------------------------------------------------|--------------------------------------------------------------------------------------------------------------------------------------------------------|----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|------------------------------------------------------------------------------------------------------------------------------|--------------------------------------------------------------------------------------------------------------------------------------------------------------------------|--------------------------------------------------------------------------------------------------------------------------------------------------------------------|------------------------------------------------------------------------------------|-------------------------------------------------------------------------------------------------------------------------------|-------------------------------------------------------------------------|-------------------------------------------------------------------------|---------------------------------------------------------------------------------------------------------------------------------------------------------------------|-----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|------------------------------------------------------------------------|
| 541 | Qy     601     ACAGTGGGGGTCTCATCCACACGTTATTAGGAATGGGACTGGACCTTCCAGGTCCTG     660       Db     585     ACAGTGGGGGTCTCATCCACACGTTATTAGGAATGGGACTGGACCTTCCAGGTCCTG     644 | ATCCC<br>                                                              |            | RESULT 7 AX032545 AX032545 AX032545 LOCUS DEFINITION Sequence 123 from Patent EP0997477. ACCESSION AX032545 VERSION AX032545. USERSION AX032545. | ISM<br>CE<br>RS                                                                                                                                        | JIILE MAC COMPLEXES and uses thereof JOURNAL Patent: EP 0997477-A 123 03-MAY-2000; SUNOL MOLECULAR CORP (US) FEATURES Location/Qualifiers Source /organism="unidentified" /mol runc="unidentified" | /db_xref="taxon:32644"                                                                                                       | Query Match 66.2%; Score 607.8; DB 2; Length 1382;<br>Best Local Similarity 89.6%; Pred. No. 1.7e-165;<br>Matches 673; Conservative 0; Mismatches 57; Indels 21; Gaps 1; | OY 1 ATGGCTCTGCAGATCCCCAGCCTCCTCTCAGCTGCTGGTGGTGGTGCTGATGGTGCTG 60                                                                                                 | Oy 61 AGCAGCCCGGGACTGAGGCGGAAACTCCATCTGCTTCTCGCCGTCGCTGGAGCACCG 120                | Oy 121 ATCGTGGTGTCCGGCAGTGGGACGGAGGTGGGGCTCATAGTGCCCCCAAGGCTCTGGA 180  Db 105 GCTGCTCACGCTGAAATCAACGAAGCTGGTCGTGCTAGCGGGGGGGG | Oy 181 GGTGGACGCTCCGAAAGGCATTTCGTGGTCCAGTTCAAGGGCGAGTGCTACTACACCAAC 240 | Oy 241 GGGACGCAGCGCATACGGCTCGTGACCAGATACATCTACAACCGGGAGGAGTACGTGCGC 300 | Oy 301 TACGACAGCGACGTGGGGGGGGTACCGCGGGGGGGGGGG                                                                                                                      | Oy 361 TACTGGAACAGCCAGAGATCCTGGAACGCGAACGCGGGCCAAGGTGGACGCGTGC 420                                                                                                                | OY 421 AGACACAACTACGAGGGCCGGAGACCAGCACCTCCCTGCGGCGGCTTGAACAGCCCAAT 480 |

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420

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999 644 720

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  Query Match 66.2%; Score 607.8; DB 2; Best Local Similarity 89.6%; Pred. No. 1.7e-165; Matches 673; Conservative 0; Mismatches 57;
  Rhode, P.R., Jiao, J.A., Burkhardt, M. and
MHC complexes and uses thereof
Patent: Ep 1556141-A 121 27-APR-2005;
Altor BioScience Corporation (US)
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  Length 1385
   Unknown.
Unclassified.
1 (bases 1 to 1385)
Rhode, P.R., Jaoo, J.-A., Burkhardt, M. and Wong, H.C.
MHC molecules and uses thereof
Patent: US 6309645-A 121 30-OCT-2001;
Location/Qualifiers
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   linear
  Score 607.8; DB 2;
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1 (bases 1 to 1508)

Rhode, P. R., Jiao, J.-A., Burkhardt, M. and Wong, H.C. Single chain MHC complexes and uses thereof

Patent: US 5868270-A 122 09-FEB-1999;

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Mer complexes and uses thereof
Patent: EP 0991477-A 121 03-MAY-2000;
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Database

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Result No.

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The specification describes soluble recombinant proteins that comprise at least a dimer formed from the alpha and beta-chains of MHC (major histocompatibility complex) Class I and II molecules in which at least one chain has, attached to its C-terminus, at least part of the Fc region of an immunoglobulin. The recombinant proteins, when linked to an antigenied, pare used to count and/or purify antigen-reactive T lymphocytes and to characterize their phenotype, e.g. in preclinical evaluation of vaccines. They are also used as immunostimulants, count and determine phenotype of autoreactive T cells in subjects with, or at risk of developing, autoimmune diseases, e.g. for staging or count and determine phenotype of autoreactive T cells in subjects with, or at risk of developing, autoimmune diseases, e.g. for staging or from cell cultures or patient samples, for use in subsequent curative or preventative cellular therapy. The present sequence encodes a recombinant protein of the invention, comprising a beta chain of MHC molecules
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  TACTOGAACAGCCGGAGATCCTGGAGCGAACGCGGGCCGAGGTGGACACGGCGTGC
   TACTGGAACAGCCAGCCGGAGATCCTGGAGCGAACGCGGGCCCGAGGTGGACACGGCGTGC
  AGACACAAACTACGAGGGGCCGGAGACCAGCACCTCCCTGCGGGGGGCTTGAACAGGCCCAAT
  GTCGCCATCTCCCTGTCCAGGACAGAGCCCTCAACCACCACACACTCTGGTCTGTTCG
   GTCGCCATCTCCCTGTCCAGGACAGAGCCCCTCAACCACCACAAAAACACTCTGGTCTGTTCG
   GTGACAGATTTCTACCCAGCCAAGATCAAAGTGCGCTGGTTCAGGAATGGCCAGGAGGAG
  GTGACAGATTTCTACCCAGCCAAGATCAAAGTGCGCTGGTTCAGGAATGGCCAGGAGGAG
   ACAGTGGGGGTCTCATCCACACATATTAGGAATGGGGACTGGACCTTCCAGGTCCTG
  ACAGTGGGGGTCTCATCCACACAGCTTATTAGGAATGGGGACTGGACCTTCCAGGTCCTG
  GTCATGCTGGAGATGACCCCTCATCAGGAGGACTCTACACCTGCCATGTGGAGCATCCC
   GTCATGCTGGAGATGACCCCTCATCAGGAGAGGTCTACACCTGCCATGTGGAGAGCATCCC
   Gaps
   ö
   Length 921;
  Sequence 921 BP; 214 A; 265 C; 286 G; 156 T; 0 U; 0 Other;
  0; Indels
  100.0%; Score 918; DB 5; L
100.0%; Pred. No. 7.3e-203;
iive 0; Mismatches 0;
  Best Local Similarity 100.
Matches 918, Conservative
   Н
   Н
  61
  241
  241
  301
   361
  421
  481
   601
  601
   721
  61
  121
  121
   181
  301
   361
   541
  661
  661
   721
   Query Match
   181
  421
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  900
   /*tag= e
/note= "binding site for primer #331 (see AAT04261)"
complement(158. .212)
  *tag= b
note= "binding site for primer #261 (see AAT04260)"
   *tag= h
'note= "probable primer binding site (primer #270)"
:11. .825
   /*tag= k
note= "probable primer binding site (primer #272)"
308. .836
   /*tag= a
/note= "probable primer binding site (primer #233)"
complement(45. .74)
   *tag= 1
note= "probable primer binding site (primer #259)"
377. .893
  note= "probable primer binding site (primer #232)"
   /note= "probable primer binding site (primer #271)"
   site"
  Polymerase chain reaction; PCR; primer; amplify; mistocompatability complex; MHC; T-cell receptor; TCR; autoimmune disease; immunodeficiency disease; immune response; immunoproliferation disease; graft-host rejection; therapy; 88.
  /*tag= 9
note= "primer #333 (see AAT04263) binding
371. .389
  "IA beta chain beta 2 region"
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/note= "primer #332 binding site"
complement(199. .250)
   "hybrid IA beta chain"
   /*tag= d
/note= "leader region"
complement(119. .172)
  Location/Qualifiers
1. 16
/*tag= a
  CATCATCATCATCAT 918
   CATCATCATCATCAT 918
   BP
   AAT04262 standard; DNA; 893
   61. 828
/*tag= c
/product= "
   /product= '
521. .550
   (first entry)
   Hybrid IA beta chain gene
  . 554
  61. .141
  *tag=
   *tag=
  16-APR-1996
  misc_feature
  WO9523814-A1
  primer_bind
   primer_bind
  sig_peptide
  primer_bind
   primer_bind
  primer_bind
  mat_peptide
  primer bind
   primer_bind
  primer_bind
  primer_bind
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   Synthetic
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ACAGTGGGGGTCTCATCCACACACATTAAGAATGGGGACTGGACCTTCCAGGTCCTG 660
                                  Greecearcreteraseacasasesecretearcreaceacacacacacacreteras
   The invention relates to a novel class II major histocompatibility compolex (MHC) antigenic peptide composite comprising a peptide containing the T-cell antigenic determinant of a mucous membrane invasive protein and the extracellular region of a class II MHC molecule or at least part of the extracellular region of the class II MHC molecule having an amino
   Peptide-Class II major histocompatibility complex (MHC) composite, useful for detecting antigen specific CD4+ T-cell, comprises antigen peptide containing epitope of mucous membrane invasive protein, and extracellular region of MHC.
  I-Ab(beta)-Cholera toxin B subunit-leucine zipper (LZ)-BirA fusion cDNA.
   flow cytometry; mucous membrane invasive antigen;
I-Ab(beta)-cholera toxin B subunit-leucine zipper-BirA fusion; CTB; ss;
  class II major histocompatibility complex; MHC; CD4+ T-cell detection;
   GTGACAGATTTCTACCCAGCCAAGATCAAAGTGCGCTGGTTCAGGAATGGCCAGGAGGAG
   GTGACAGATTTCTACCCAGCCAAGATCAAAGTGCGCTGGTTCAGGAATGGCCCAGGAGAG
  ACAGTGGGGGTCTCCACACACCTTATTAGGAATGGGGACTGGACCTTCCAGGTCCTG
   GTCATGCTGGAGATGACCCCTCATCAGGGAGGTCTACACCTGCCATGTGGAGCATCCC
   706 Greatscreakardaccerteareassaaassarcraecersecarsresassarcee
   AGCCTGAAGAGCCCCATCACTGTGGAGTGGAGGCACAGTCCGAGTCTGCCCGGAGCAAG
  766 AGCCTGAAGAGCCCCATCACTGTGGAGTGGAGGGCACAGTCCGAGTCTGCCCGGAGCAAG
/product= "I-Ab(beta)-Cholera toxin B subunit (CTB)-
leucine zipper (LZ)-BirA fusion cDNA"
  (SENT-) SENTAN KAGAKU GLJUTSU INCUBATION CENT KK
   Example 1; SEQ ID NO 10; 30pp; Japanese.
  Location/Qualifiers
  BP.
   ADO31225 standard; cDNA; 945
   03-DEC-2003; 2003JP-00404367.
   03-DEC-2002; 2002JP-00351818
  (first entry)
   ๗
   GGAGGTGGAGG 791
  826 TAAGCATGCGG 836
  1. .945
/*tag=
  WPI; 2004-546819/53.
P-PSDB; ADQ31224.
   Vibrio cholerae.
Unidentified.
   JP2004196789-A
  07-OCT-2004
   646
  586
   ADQ31225;
   541
   601
   661
  721
  gene.
  Key
   ADQ31225
   RESULT
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  7
   This sequence represents a hybrid IA beta chain gene, containing the chicken ovalbumin peptide (COVA). This sequence was used in the construction of a hybrid IA alpha beta dimer. The encoded protein (plAd-OVA) was found to be more stable than the IA alpha beta dimer. The stability was decreased by the addition of a MHC groove specific binding peptide (e.g. see ARR82527, AAR82528 and AAR82511), compared to an increase seen on the addition of a MHC binding peptide to IE k/d-MCC. These complexes may be used to regulate an immune response. The complexes are capable of being recognised by a TCR alone or in combination with additional MHC proteins. These complexes are useful for therapeutic additional MHC proteins. These complexes are useful for therapeutic purposes and experimental purposes. They can also be used as reagents for the treatment of diseases including autoimmune diseases, immunodeficiency diseases, immunodeficiency
   240
  300
  345
   360
  405
  480
   525
   540
   Arescrerecasareceasecreciererereseristicasers and a secretaristic secret
   420
  AGCAGCCCCGGGACTGAGGGCGGAAACTCCATCTGCTTCTCGCCGTCGCTGGAGCACCCG 120
  Peptide-MHC complex comprising antigenic peptide, linker and MHC segment - useful as reagents for the treatment of diseases including auto-immune diseases, immuno-stimulatory diseases or graft-host rejection.
  9
  ATGAGGCTGCCAG------AGGAGGTGGGGCTCACTAGTGCCCCGAGGCTCTGGA
  GGTGGGAGGCTCCGAAAGGCATTTCGTGGTCCAGTTCAAGGGCGAGTGCTACTACACCAAC
   1 ATGGCTCTGCAGATCCCCAGCCTCCTCTCAGCTGCTGTGGTGGTGGTGATGGTGCTG
   ATCGTGGTGTCCGGCAGCTGGGACGTGGGGGCTCACTAGTGCCCCCGAGGCTCTGGA
  GGTGGAGGCTCCGAAAGGCATTTCGTGGTCCAGTTCAAGGGCGAGTGCTACTACAAC
  GGGACGCAGCGCATACGGCTCGTGACCAGATACATCTACAACCGGGAGGAGTACGTGCGC
   GGGACGCAGCGCATACGCCTCGTGACCAGATACATCTACAACCGGGAGGAGTACGTGCGC
   TACGACAGGGACGTGGGCGAGTACCGCGGGGGGCGGGCTGGGGCGGCCCAGACGCCGAG
  TACTGGAACAGCCGGAGATCCTGGAGCGAACGCGGGCCGAGGTGGACACGGCGTGC
  AGACACAACTACGAGGGCCGGAGACCAGCACCTCCCTGCGGCGGCTTGAACAGCCCAAT
   GTCGCCATCTCCCTGTCCAGGACAGAGCCCTCAACCACCACAACACTCTGGTTCG
  Gaps
  15;
  Length 893;
   Sequence 893 BP; 204 A; 239 C; 275 G; 175 T; 0 U; 0 Other;
  Indels
  Score 696.2; DB 2;
Pred. No. 1.9e-151;
0; Mismatches 28;
   (NAJE-) NAT JEWISH CENT IMMUNOLOGY & RESPIRATORY
   Example 1; Page 53; 94pp; English
  95WO-US002689
  94US-00207481
  ch
11 Similarity 94.6%;
748; Conservative
  Marrack P;
   WPI; 1995-320543/41.
  Local Similarity
   P-PSDB; AAR82533.
  03-MAR-1995;
  04-MAR-1994;
                            08-SEP-1995
   Kappler JW,
  61
  61
   121
  176
   286
  346
   406
  481
  Query Match
  121
  226
   361
   421
  181
  241
  301
   Matches
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acid sequence comprising one or more deletions, substitutions or additions. The molecule of the invention may be useful for detecting an antigen-specific CD4+ T-cell by flow cytometry and for presenting an microorganism-derived mucous membrane invasive protein as an antigen. The method of the invention enables efficient detection of antigen-specific activation of CD4+ T-cells in the mucous membrane. The current sequence is that of the class II major histocompatibility complex-related I-Ab(alpha)-Cholera toxin B subunit (CTB)-leucine zipper (LZ)-BirA fusion
   282
   342
   405
   522
   762
   225
  345
   402
   462
   525
   582
   585
   642
   645
   702
   705
   765
   819
   882
   879
  Greacacreáregrecreadecrececacreacimmederegaderecerecereceaae 108
   GAGTGCTACTTCACCAACGGGACGCAGCGCATACGATATGTGACCAGATACATCTACAAC
   CGGGAGGAGTACGTGCGCTACGACACGACGTGGGCGAGCACCACCGCGCGGTGACCGAGCTG
   GGGCGGCCAGACGCCGAGTACTGGAACAGCCAGCCGGAGATCCTGGAGCGAACGCGGGCC
  CGGCTTGAACAGCCCAATGTCGTCATCTCCCTGTCCAGGACAGAGGCCCTCAACCACCAC
  TGGACCTTCCAGGTCCTGGTCATGCTGGAGATGACCCCTCGGCGGGGAGAGGTCTACACC
   AAGAAACTGCAGGCACTTAAGAAAAAGAACGCTCAGCTGAAGTGGAAACTTCAAGCCCTC
  GTGGTGCTGATGGTGCTGAGCAGCCCCGGGACTGAGGGCGGAAACTCCATCTGCTTCTCG
  CCGTCGCTGGAGCACCCGATCGTGGTGTCCCGGCAGCTGGGACGTGGGGGGCTCACTA
  GTGCCCCGAGGCTCTGGAGGTGGAGGCTCCGAAAGGCATTTCGTGGTCCAGTTCAAGGGC
   GCAGGGGGAAG---TGGAGGTGGAGGGTCTGAAAGGCATTTCGTGTACCAGTTCATGGGC
   GAGTGCTACTACACCCAACGGGACGCAGCGCATACGGCTCGTGACCAGATACATCTACAAC
   CGGGAGGAGTACGTGCGCTACGACACGCGACGTGGCCGAGTACCGCGGTGACCGAGCTG
  GGGCGGCCAGACGCCGAGTACTGGAACAGCCAGCCGGAGATCCTGGAGCGAACGCGGGCC
  GAGGTGGACACGCGTGCAGACACAACTACGAGGGCCGGAGACCAGCACCTCCCTGCGG
   CGGCTTGAACAGCCCAATGTCGCCATCTCCCTGTCCAGGACAGAGGCCCTCAACCACCAC
  AACACTCTGGTCTGTTCGGTGACAGATTTCTACCCAGCCAAGATCAAAGTGCGCTGGTTC
   TGGACCTTCCAGGTCCTGGTCATGCTGGAGATGACCCCTCATCAGGGAGGTCTACACC
  TECCATETEGAGCATCCCAGCCTGAAGAGCCCCCATCACTGTGGAGTGGAGGGCCACAGTCC
  GAGTCTGCCCGGAGCAAGGGAGGTGGAGGATCCACTACAGCTCCATCAGCTCAGTTGAAA
  AAGAAATTGCAAGCACTGAAGAAAAAAAGGCTCAGCTGAAGTGGAAACTTCAAGCCCTC
  Gaps
  ٠.
د
  Length 945;
  Sequence 945 BP; 230 A; 256 C; 294 G; 165 T; 0 U; 0 Other;
  Score 665.2; DB 12; Length
Pred. No. 3e-144;
0; Mismatches 108; Indels
  72.5%;
  759; Conservative
  cDNA of the invention.
   Query Match
Best Local Similarity
   994
  43
   49
   103
   109
  163
  169
   223
   283
  286
  343
   346
  403
   406
   463
   466
  523
  526
   583
  586
  643
  646
  703
   904
  763
  823
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883 AAGAAGAAACTCGCCCAGCATCATCATCATCAT 918

```
for primer #362 (see AAT04270)"
   primer #363 (see AAT04271)"
   Major histocompatability complex; MHC; T-cell receptor; TCR; autoimmune disease; immunodeficiency disease; immune response; immunoproliferation disease; graft-host rejection; therapy; B cell; M12.C3; pM12-IAb-Ea; ss.
  note= "probable primer binding site (primer #271)"
44. .568
  site (primer #272)"
  site (primer #259)"
  #270) "
   #366) "
  /*tag= n
/note= "probable primer binding site (primer #59)"
   /*tag= a
/note= "probable primer binding site (primer #76)"
complement(40. .74)
   aite"
  *tag= h
note= "probable primer binding site (primer
31. .959
   (primer
  *tag= g
'note= "primer #365 (see AAT04272) binding
185. .403
  beta chain beta 2 region"
   site
                         915
  /*tag= f
/note= "primer #364 binding site"
complement(212. .266)
  chain"
   *trag= k
note= "probable primer binding
823. .850
  *tag= m
note= "probable primer binding
.000. .1013
  primer binding
AAGAAGAAACTCGCCCAGCTGCATCATATTCTGGAT
  "hybrid IA beta
   63. .143
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/note= "leader region"
   /*tag= b
/note= "binding site
  Location/Qualifiers
  note= "probable
  BP.
  "IA
  95WO-US002689
  DNA; 1013
   94US-00207481
   /*tag= c
/product= '
  product=
  (first entry)
   Hybrid IA beta chain gene
   535. .564
  63. .959
   *tag=
   *tag=
   . .18
*tag≕
   *tag=
  standard;
  WO9523814-A1
   04-MAR-1994;
  03-MAR-1995;
  16-APR-1996
  primer_bind
  primer_bind
   primer_bind
   primer_bind
   primer_bind
  sig_peptide
  primer_bind
   primer_bind
   mat_peptide
   primer_bind
  primer_bind
  primer bind
  primer_bind
   Synthetic
                         880
  AAT04269
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WPI; 2004-546819/53.
  coli.
   P-PSDB; ADQ31227
  Sequence 915 BP;
  JP2004196789-A
   07-0CT-2004
  Escherichia
Unidentified
   region of
  721
   ADQ31228;
                                 661
  ss; gene.
   Key
   셤
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   g
  1;
   This sequence represents a hybrid IA beta chain gene. This sequence contains a fragment of the IE alpha chain (residues 56-73), as well as a linker and cleavage site. This sequence was transfected into a B cell line (M12.C3) using plasmid pM12-IAb-Ba. It was found that the encoded sequence was expressed in these cells. Complexes such as this may be used to regulate an immune response. The complexes are capable of being recognised by a TCR alone or in combination with additional MHC proteins. These complexes are useful for therapeutic purposes and experimental purposes. They can also be used as reasgents for the treatment of diseases including autoimmune diseases, immunodeficiency diseases.
   419
   420
   479
   629
   122
   182
   180
  239
   240
   300
   359
  360
   480
   539
   540
   599
  600
   099
  AGCAGCCCCGGGACTGAGGCGGAAACTCCATCTGCTTCTCGCCGTCGCTGGAGCACCCG 120
  Peptide-MHC complex comprising antigenic peptide, linker and MHC segment - useful as reagents for the treatment of diseases including auto-immune diseases, immuno-stimulatory diseases or graft-host rejection.
   9
   Areacreracanteceascerecreereresereresereseresereseres
  GGTGGAGGGTCTGAAAGGCATTTCGTGTACCAGTTCATGGGCGAGTGCTACTTCACCAAC
  GGGACGCACGCATACGATATGTGACCAGATACATCTACAACCGGGAGGAGTACGTGCGC
   1 ATGGCTCTGCAGATCCCCAGCCTCCTCTCAGCTGCTGTGGTGGTGGTGGTGGTGCTG
  ATCGTGGTGTCCGGCAGCTGGGAGGTGGGGCCTCACTAGTGCCCCCGAGGCTCTGGA
   GCCAACATTGCTGTCGACAAGCTGGAGGTGGATC---CGGTGGAGGGGGAAGTGGA
  GGTGGAGGCTCCGAAAGGCATTTCGTGGTCCAGTTCAAGGGCGAGTGCTACTACACCAAC
   GGGACGCAGCGCATACGCCTCGTGACCAGATACATCTACAACCGGGAGGAGTACGTGCGC
  TACTGGAACAGCCAGCCGGAGATCCTGGAGCGAACGCGGGCCGAGGTGGACACGGCGTGC
   AGACACAACTACGAGGGCCGGAGACCAGCACCTCCCTGCGGCGCGCTTGAACAGCCCAAT
   AGACACAACTACGAGGGGCGGGGGGGCCCCACCCCCCCGGGGGGCGCTTGAACAGCCCAAT
  GTCGTCATCTCCCTGTCCAGGACGCCCTCAACCACCACACACTCTGGTCTCA
  GTGACAGATTTCTACCCAGCCAAGATCAAAGTGCGCTGGTTCAGGAATGGCCAGGAGGAG
  GTGACAGATTTCTACCCAGCCAAGATCAAAGTGCGCTGGTTCCCGGAATGGCCAGGAGGAG
   Gaps
   .;
m
   Length 1013;
   Sequence 1013 BP; 220 A; 272 C; 327 G; 192 T; 0 U; 2 Other;
  Indels
   immunoproliferation diseases, and graft-host rejection
   81;
 IMMUNOLOGY & RESPIRATORY
  Score 634; DB 2; I Pred. No. 5.1e-137;
  1; Mismatches
  Example 2; Page 65; 94pp; English
  Query Match
Best Local Similarity 89.1%;
Matches 695; Conservative
 JEWISH CENT
                           Marrack P;
   WPI; 1995-320543/41.
P-PSDB; AAR82538.
NAT
                         Kappler JW,
  241
  63
  61
  123
   121
   183
  181
  240
   300
  301
  360
  361
   420
  421
  480
  481
   540
  009
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  601
(NAJE-)
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The invention relates to a novel class II major histocompatibility complex (MHC) antigenic peptide composite comprising a peptide containing the T-cell antigenic determinant of a mucous membrane invasive protein and the extracellular region of a class II MHC molecule or at least part of the extracellular region of the class II MHC molecule having an amino acid sequence comprising one or more deletions, substitutions or additions. The molecule of the invention may be useful for detecting an antigen-specific CD4+ T-cell by flow cytometry and for presenting a microorganism-derived mucous membrane invasive protein as an antigen. The method of the invention enables efficient detection of antigen-specific activation of CD4+ T-cells in the mucous membrane. The current sequence is that of the class II major histocompatibility complex-related I-Ab(alpha)-Escherichia coli heat-labile toxin B subunit (LTB)-leucine
   780
  660 ACGGTGGGGCTCTCATCCACACAGCTTATTAGGAATGGGGACTGGACCTTCCAGGTCCTG 719
  Peptide-Class II major histocompatibility complex (WHC) composite, useful for detecting antigen specific CD4+ T-cell, comprises antigen peptide containing epitope of mucous membrane invasive protein, and extracellular region of MHC.
  /product= "I-Ab(beta)-Escherichia coli heat-labile toxin
B subunit (LTB)-leucine zipper (LZ)-BirA fusion protein"
   flow cytometry; mucous membrane invasive antigen;
I-Ab(beta)-heat-labile toxin B subunit-leucine zipper-BirA fusion; LTB;
   class II major histocompatibility complex; MHC; CD4+ T-cell detection;
   720 GTCATGCTGGAGATGACCCCTCGGCGGGGAGGTCTAYACCTGTCACGTGGAGCATCCC
  AGCCTGAAGAGCCCCCATCACTGTGGAGTGGAGGGCACAGTCCGAGTCTGCCCGGAGCAAG
  I-Ab(beta)-E. coli heat-labile toxin B subunit-LZ-BirA fusion cDNA.
  Length 915;
   228 A; 242 C; 271 G; 174 T; 0 U; 0 Other;
  DB 12;
   --
  (SENT-) SENTAN KAGAKU GIJUTSU INCUBATION CENT KK
   zipper (LZ)-BirA fusion cDNA of the invention.
  Score 628.6;
  Example 3; SEQ ID NO 13; 30pp; Japanese.
  Location/Qualifiers
1. .915
/*tag= a
   BP.
   ADQ31228 standard; cDNA; 915
  03-DEC-2003; 2003JP-00404367.
  03-DEC-2002; 2002JP-00351818
  68.5%;
  (first entry)
  Query Match
  ACAGTGGGGGTCTCATCCACACAGCTTATTAGGAATGGGGACTGGACCTTCCAGGTCCTG
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  421
   394
  481
   454
  541
   514
   601
  574
  661
  634
  721
  694
  781
  901
   TGGTCATCTCCCTGTCCAGGACAGAGGCCCTCAACCACCACAACACTCTGGTCTGCTCAG
  TGACAGATTTCTACCCAGCCAAGATCAAAGTGCGCTGGTTCCGGAATGGCCAGGAGAA
   TTCCGCGCGGAATCCACTACAGCTCCATCAGCTCAGTTGAAAAAGAAAACTGCAGGCACTTA
   AGAAAAAGAACGCTCAGCTGAAGTGGAAACTTCAAGCCCTCAAGAAGAAACTCGCCCAGC
                   GTGGAGGCTCCGAAAGGCATTTCGTGGTCCAGTTCAAGGGCGAGTGCTACTACACCAACG
                          GGACGCAGCGCATACGGCTCGTGACCAGATACATCTACAACCGGGAGGAGTACGTGCGCT
  ACGACAGGGACGTGGGCGGACCGCGGGGTGACCGAGCTGGGGGCGGCCAGACGCCGAGT
  ACTGGAACAGCCAGCCGGAGATCCTGGAGCGAACGCGGGCCGAGGTGGACACGGCGTGCA
   TGACAGATTTCTACCCAGCCAAGATCAAAGTGCGCTGGTTCAGGAATGGCCAGGAGGAGA
  CAGTGGGGGTCTCATCCACACAGCTTATTAGGAATGGGGACTGGACCTTCCAGGTCCTGG
   TCATGCTGGAGATGACCCCTCATCAGGGAGGTCTACACCTGCCATGTGGAGCATCCCA
   GCCTGAAGAGCCCCATCACTGTGGAGTGGAGGGCACAGTCCGGAGTCTGCCCGGAGCAAGG
  GAGGIGGAGGAICCACTACAGCICCATCAGCICAGTIGAAAAAGAAATIGCAAGCACTGA
      Gaps
  Construction; major histocompatibility complex; MHC; fusion complex;
       .
9
      Indels
8.9e-136;
thes 54;
Pred. No. 8.96
); Mismatches
   Location/Qualifiers 6. .1382
  BP
      ö
   ATCATCATCATCAT 918
  AAT86989 standard; DNA; 1382
91.94;
   TGCATCATATTCTGGAT
   SCEl single chain gene; ss
       Conservative
  SCEl single chain gene
  (first
Similarity
   27-MAR-1998
   242
   215
   275
  362
  335
  422
   395
   782
  749
  842
   902
   698
                   182
                              155
   302
  482
   455
  542
  515
  575
   662
   635
  722
   AAT86989
  602
  695
Best Local
Matches 67
   Key
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180
  Single chain major histocompatibility complex comprising linked alpha and beta chains - useful for suppressing an immune response to an auto:immune disease, e.g. multiple sclerosis, rheumatoid arthritis, diabetes mellitus, etc.
   584
   AGCAGCCCCGGGACTGAGGGCGGAAACTCCATCTGCTTCTCGCCGTCGCTGGAGCACCCG 120
   164
   300
   360
   344
  420
   404
   TACGACAGCGACGAGGCGAGTACCGCGGGGGGACCGAGCTGGGGCGGCGCAAGCCGAG
  GTCGCCATCTCCCTGTCCAGGACAGAGGCCCTCAACCACCACACACTCTGGTCTGTTCG
   AGCAGCCCAAGGAC-------AGCTAAGTATCTCTCAGGCTGTTCAC
   gerecreacecreaarcaaceáaecreercerecraecesaeceaeceaaececeaa
  eggaccagcarracgccrcgrgaccagaracarcracaaccgggaggaggagaracgcgc
   TACGACACGACGTGGGCGAGTACCGCGGTGACCGAGCTGGGGCGCCCAGACGCCGAG
  TACTGGAACAGCCAGCCGGAGATCCTGGAGCGAACGCGGGCCGAGGTGGACACGGCGTGC
   AGACACAACTACGAGGGCCGGAGACCAGCACCTCCCTGCGGCGGCTTGAACAGCCCAAT
   ATGGCTCTGCAGATCCCCAGCCTCCTCTCAGCTGCTGTGGTGGTGCTGATGGTGCTG
  121 ATCGTGGTGTCCGGCAGCTGGGAGGTGGGGGCTCACTAGTGCCCCGAGGCTCTGGA
  GGTGGAGGCTCCGAAAGGCATTTCGTGGTCCAGTTCAAGGGCGAGTGCTACTACACCAAC
   GGGACGCAGCGCATACGGCTCGTGACCAGATACATCTACAACCGGGAGGAGTACGTGCGC
   Gaps
   21;
  Length 1382;
  Sequence 1382 BP; 320 A; 373 C; 405 G; 284 T; 0 U; 0 Other;
  of major
   Indels
  The present sequence was used in the construction histocompatibility complex (MHC) fusion complexes
   57;
  DB 2;
  Score 607.8; DB 2;
Pred. No. 6.5e-131;
0; Mismatches 57;
  HC;
   Example 17; Page 140-141; 217pp; English
  Wong
  Σ
  Burkhardt
  66.2%;
89.6%;
  97WO-US001617
  96US-00596387
 σ
  Query Match
Best Local Similarity 89.6'
Matches 673; Conservative
/*tag=
   (DADE-) DADE INT INC
  WPI; 1997-402555/37.
  Jiao J,
   P-PSDB; AAW29214.
  31-JAN-1996;
  30-JAN-1997;
                                  W09728191-A1
   07-AUG-1997
  PR,
  405
   285
   361
  345
   99
   181
   241
   225
  301
  541
```

104

9 9 240 224 284

beta chains of the MHC fusion complex.

```
The invention relates to a major histocompatibility complex (WHC) fusion complex (I) comprising an WHC molecule that contains a peptide-binding 20 groove, and a presenting peptide covalently (e.g. an antigenic peptide) linked to the MHC molecule, where (I) is capable of modulating the activity of a T cell. Also included are a DNA construct coding for the complex, where the MHC molecule is a class II MHC (e.g. mouse I-Ad or I-As, or human HLA-DRI (human leukocyte antigen-DRI)), a multivalent MHC tusion complex comprising two or more linked complexes, identifying a peptide that can modulate the activity of T cells (involving introducing into host cells cloning vectors that each contain the fusion complex, and selecting host cells that express MHC fusion complex, and selecting host cells that express MHC fusion complex, and selecting host cells that and beta combinant expression vector comprising DNA that codes for the alpha and beta chains of the fusion complex MHC protein, a single recombinant expression vector comprising DNA that codes for a T cell costimulatory factor and the alpha
                                       644
  720
  704
  Novel major histocompatibility complex fusion complex having presenting peptide covalently linked to MHC molecule containing peptide-binding groove, used for suppressing immune response in multiple sclerosis,
 ACAGTGGGGGTCTCATCCACACAGCTTATTAGGAATGGGGACTGGACCTTCCAGGTCCTG
                               ACAGTGGGGGTCTCATCCACACACTTATTAGGAATGGGGACTGGACCTTCCAGGTCCTG
  Greatgerdandardaccentearcadadadadereraceraceargadaderece
  GTCATGCTGGAGATGACCCCTCATCAGGGAGGTCTACACCTGCCATGTGGAGCATCCC
   MHC; major histocompatibility complex; gene therapy; fusion complex; peptide-binding groove; T cell modulation; class II MHC; vaccine; autoimmune disorder; multiple sclerosis; rheumatoid arthritis; insulin-dependent diabetes mellitus; myasthenia gravis; immunogen; chronic allergy; mouse; ds; I-Ad; gene.
   S, Edwards AC;
   Mouse MHC I-Ad/Ova 323-339 synthetic gene SCE1.
  AGCCTGAAGAGCCCCATCACTGTGGAGTGGA 735
   751
   Weidanz JA, Grammer
  AGCCTGAAGAGCCCCATCACTGTGGAGTGGA
   Example 17; Fig 29; 126pp; English
   BP.
   ACA60744 standard; DNA; 1382
  94US-00283302.
   2001US-00900379
   95US-00382454
97US-00776084
  16-JUN-2003 (first entry)
   Jiao JJJ;
   DADE-) DADE INT INC.
   2003-341126/32.
   Rhode PR,
  P-PSDB; ABU72108
  US2002198144-A1.
   Chavaillaz P,
   29-JUL-1994;
01-FEB-1995;
17-JAN-1997;
   06-JUL-2001;
   26-DEC-2002
   Mus sp.
Synthetic.
  645
  allergies
                                     585
601
   661
  721
  ACA60744;
   HC,
   Wong
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The expression of the fusion complex. Also included are inducing an immune response in a mammal all sequences and Kozak sequence for efficient expression of the fusion complex. Also included are inducing an immune response in a mammal against a targeted disorder, by administering DNA sequence comprising a fusion complex, or DNA sequence coding for a fusion complex which is a single chain fusion molecule) and suppressing an immune response in a mammal by administering to the mammal a DNA sequence comprising a transmembrane domain, and a contenting peptide that contains a transmembrane domain, and a consist and is covalently linked to the MHC protein, or DNA sequence coding for the fusion complex which is a single chain fusion molecule. The methods are useful for identifying a peptide that can modulate the cativity of T cells, inducing an immune response in a mammal (including vaccinating a mammal against a targeted disorder) and for suppressing an immune response in a mammal include an autoimmune disorder such as multiple sclerosis, insulin-dependent diabetes mellitus, the present sequence encodes a mouse MHC class II I-Ad fusion complex of the
  300
  284
  360
  420
  480
  344
   540
  65
  404
   464
   524
   600
   584
   999
   644
   GTCATGCTGGAGATGACCCCTCATCAGGGAGAGTCTACACCTGCCATGTGGAGCATCCC 720
   GGGACGCAGCGCATACGGCTCGTGACCCAGATACATCTACAACCGGGAGGAGGAGTACGTGCC
  61 AGCAGCCCCGGGACTGAGGGCGGAAACTCCATCTGCTTCTCGCCGTCGCTGGAGCACCCG
  GGTGGAGGCTCCGAAAGGCATTTCGTGGTCCAGTTCAAGGGCGAGTGCTACTACACCAAC
  GGGACGCAGCGCATACGGCTCGTGACCAGATACATCTACAACCGGGAGGAGTACGTGCGC
  TACGACACGACGTGGGCGAGTACCGCGCGGGGGCGAGCTGGGGCGGCCAGACGCCGAG
   ACAGTGGGGGTCTCATCCACACGTTATTAGGAATGGGGACTGGACCTTCCAGGTCTG
   1 ATGGCTCTGCAGATCCCCAGCCTCCTCTCAGCTGCTGTGGTGGTGCTGATGGTGCTG
  6 Argerrergeagareceageerecreereracracracracracracracracracracracra
  121 ATCGTGGTGCCGCCAGCTGGACGCAGGTGGGGGCTCACTAGTGCCCCGAGGCTCTGGA
  TACTGGAACAGCCAGCCGGAGATCCTGGAGCGAACGCGGGCCGAGGTGGACACGGCGTGC
   TACTGGAACAGCCAGCCGGAGTCCTGGAGCGAACGCGGGCCGAGGTGGACACGGCGTGC
   AGACACAACTACGAGGGCCGGAGACCAGCACCTCCCTGCGGCGGCGGCTTGAACAGCCCAAT
   AGACACAACTACGAGGGCCCGGAGACCAGCACCTCCCTGCGGCGCCTTGAACAGCCCAAT
   GTCGCCATCTCCCTGTCCAGGACAGAGCCCTCAACCACCACACACTCTGGTCTGTTCG
   GTCGCCATCTCCCTGTCCAGGACAGACCCTCAACCACCACCACCACTCTGGTCTGTTCG
  GTGACAGATTTCTACCCAGCCAAGATCAAAGTGCGCTGGTTCAGGAATGGCCAGGAGGAG
   GTGACAGATTTCTACCCAGCCAAGATCAAAGTGCGCTGGTTCAGGAATGGCCAGGAGGAG
   ACAGTGGGGGTCTCATCCACACACACTTATTAGGAATGGGGACTGGACCTTCCAGGTCCTG
   Gaps
   21;
   Length 1382;
   Sequence 1382 BP; 320 A; 373 C; 405 G; 284 T; 0 U; 0 Other;
   Indels
   DB 8;
  .5e-131;
les 57;
  Pred. No. 6.5e
0; Mismatches
   Score 607.8;
Pred. No. 6.5
   ch
11 Similarity 89.6%;
673; Conservative
   Invention
  105
   Query Match
Best Local S
  181
  165
  241
  225
  285
  361
   465
   525
  301
  345
   405
   601
   585
   421
   661
   481
   541
   Matches
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   404
   480
  720
 300
                  284
   420
  464
   540
   524
  900
   584
   999
   644
   704
           GGGACGCAGCGCATACGGCTCGTGACCAGATACATCTACAACCGGGAGGAGTACGTGCGC
  GTCGCCATCTCCCTGTCCAGGACAGAGGCCCTCAACCACCACAAACACTCTGGTCTGTTCG
  GICATGCTGGAGATGACCCCTCATCAGGAGAGGTCTACACCTGCCATGTGGAGCATCCC
   AGACACAACTACGAGGGCCGGAGACCAGCACCTCCCTGCGGCGCCTTGAACAGCCCAAT
  GTGACAGATTTCTACCCAGCCAAGATCAAAGTGCGCTGGTTCAGGAATGGCCAGGAGGAG
  601 ACAGTGGGGGTCTCATCCACACATATAGGAATGGGGACTGGACCTTCCAGGTCCTG
   Acadregesercricarceacacerrarrassaaresesacresacerrecassicers
   GTCATGCTGGAGATGACCCCTCATCAGGGAGGTCTACACCCTGCCATGTGGAGCATCCC
GGGACGCAGCGCATACGGCTCGTGACCAGATACATCTACAACCGGGAGGAGTACGTGCGC
   TACTGGAACAGCCAGCCGGAGATCCTGGAGCGAACGCGGGCCCGAGGTGGACACGCCGTGC
  MHC; major histocompatibility complex; gene therapy; fusion complex; peptide-binding groove; T cell modulation; class II MHC; vaccine; autoformune disorder; multiple sclerosis; rheumatoid arthritis; insulin-dependent diabetes mellitus; myasthenia gravis; immunogen; chronic allergy; mouse; ds; I-Ad; gene.
   Edwards AC;
   Mouse MHC I-Ad/Ova 323-339 synthetic gene SSCI.
  Grammer S,
  AGCCTGAAGAGCCCCATCACTGTGGAGTGGA 751
   AGCCTGAAGAGCCCCATCACTGTGGAGTGGA 735
  Weidanz JA,
   BP.
  94US-00283302.
95US-00382454.
97US-00776084.
   ACA60742 standard; DNA; 1385
  06-JUL-2001; 2001US-00900379
  16-JUN-2003 (first entry)
  Chavaillaz P, Jiao JJJ;
  (DADE-) DADE INT INC
  Rhode PR,
   WPI; 2003-341126/32.
   P-PSDB; ABU72106
   US2002198144-A1.
   29-JUL-1994;
01-FEB-1995;
17-JAN-1997;
   26-DEC-2002
  Mus sp.
Synthetic.
   241
                       225
   301
  285
  361
   421
   481
   465
   541
   525
   585
   661
  645
   721
  705
   ACA60742;
  Wong HC,
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  180
  Single chain major histocompatibility complex comprising linked alpha and beta chains - useful for suppressing an immune response to an auto:immune disease, e.g. multiple sclerosis, rheumatoid arthritis, diabetes mellitus, etc.
  AGCAGCCCAAGGAC------CTTAAGTATCTCTCAGGCTGTTCAC 104
  164
   GGTGGAGGCTCCGAAAAGGCATTTCGTGGTCCAGTTCAAGGGCGAGTGCTACTACACCAAC 240
  224
  65
   9
   GTCATGCTGGAGATGACCCCTCATCAGGGAGAGGTCTACACCTGCCATGTGGAGCATCCC 704
  AGCAGCCCCGGGACTGAGGGCGGAAACTCCATCTGCTTCTCGCCCGTCGCTGGAGCACCCG
  1 ATGGCTCTGCAGATCCCCAGCCTCCTCTCAGCTGCTGCTGGGTGGTGCTGATGGTGCTG
  ATCGTGGTGTCCGGCAGCTGGGACGAGGTGGGGGCTCACTAGTGCCCCGAGGCTCTGGA
  gergercacgergaaarcaacgaagergergeraccaaggaggaggagggagggagg
   Gaps
  Construction, major histocompatibility complex; MHC; fusion complex;
  21;
   Query Match 66.2%; Score 607.8; DB 2; Length 1385; Best Local Similarity 89.6%; Pred. No. 6.5e-131; Matches 673; Conservative 0; Mismatches 57; Indels 21;
   Sequence 1385 BP; 316 A; 383 C; 399 G; 287 T; 0 U; 0 Other;
   The present sequence was used in the construction of major histocompatibility complex (MHC) fusion complexes
                                   751
  HC;
  Example 17; Page 135-137; 217pp; English.
  Wong
   Location/Qualifiers
6. .1385
/*tag= a
  Σ
   BP
  Jiao J, Burkhardt
   AAT86987 standard; DNA; 1385
   97WO-US001617,
  96US-00596387
   SSC1 single chain gene; ss
  (first entry)
   SSC1 single chain gene
  INT INC.
  WPI; 1997-402555/37.
   P-PSDB; AAW29212
   (DADE-) DADE
  30-JAN-1997;
  31-JAN-1996;
  WO9728191-A1
  27-MAR-1998
  07-AUG-1997
  Rhode PR,
  Synthetic
   705
   645
  9
  61
   99
   121
  105
  181
                         721
  AAT86987;
   Key
  œ
  RESULT
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Novel major histocompatibility complex fusion complex having presenting peptide covalently linked to MHC molecule containing peptide-binding groove, used for suppressing immune response in multiple sclerosis, allergies

Fig 27; 126pp; English. Example 17;

The invention relates to a major histocompatibility complex (WHC) fusion complex (I) comprising an MHC molecule that contains a peptide-binding of complex (I) comprising an MHC molecule that contains a peptide-binding of groove, and a presenting peptide covalently (e.g. an antigenic peptide) linked to the MHC molecule, where (I) is capable of modulating the cativity of a Teall. Also included are a DNA construct coding for the complex, where the MHC molecule is a class II MHC (e.g. mouse I-Ad or I-As, or human HLA-DR1 (human leukocyte antigen-DR1), a multivalent MHC (fusian complex, comprising two or more linked complexes, identifying a peptide that can modulate the activity of T cells (involving introducing into host cells ander conditions usitable for expression of the complex that modulate the activity of T cells (involving introducing into host complex, and selecting host cells that express MHC fusion complex, and selecting host cells that expression of the cappession vector comprising DNA that codes for a realls and beta chains of the fusion complex MHC protein, a single recombinant expression vector comprising DNA that codes for a T cell costimulatory factor and the alpha and beta chains of the MHC fusion complex. Also included are inducing an immune response in a mammal (including vaccinating a mammal against a targeted disorder, by administering DNA sequence comprising a mammal and suppressing an immune response in a mammal and suppressing an immune response in a mammal and suppressing an immune response in a mammal and suppressing an immune response in a mammal and suppressing an immune response in a mammal and suppressing an immune response in a mammal and suppressing an immune response comprising an expression vector, encoding a general peptide that is a T cell receptor (TCR) antagonist or partial against the fusion complex which is a single chain fusion modeled are the mammal and suppressing an immune response in a mammal including an immune response in a mammal including an immune response in a mamm immune response in a mammal. The disorders include an autoimmune disorder such as multiple sclerosis, insulin-dependent diabetes mellitus, rheumatoid arthritis, myasthenia gravis or chronic allergies. The present sequence encodes a mouse MHC class II I-Ad fusion complex of the

Sequence 1385 BP; 316 A; 383 C; 399 G; 287 T; 0 U; 0 Other;

240 AGCAGCCCCGGGACTGAGGGCGGAAACTCCATCTGCTTCTCGCCGTCGCTGGAGCACCCG 120 164 224 284 TACGACAGCGACGAGGGAGTACCGCGGGGGAGCCGAGCTGGGGCGGCCAGACGCCGAG 360 9 9 GGGGGAAACTCCGAAAGGCATTTCGTGGTCCAGTTCAAGGGCGAGTGCTACTACAAC 1 ATGCCTCTGCAGATCCCCAGCCTCCTCTCAGCTGCTGTGGTGGTGGTGGTGGTGCTG 121 ATCGTGGTGTCCGGCAGCTGGGACGTGGGGGCTCACTAGTGCCCCGAGGCTCTGGA GCTGCTCACGCTGAAATCAACGAAGCTGGTCGTGCTAGCGGAGGGGGCGGAAGCGGCGGA GGTGGAGGCTCCGAAAGGCATTTCGTGGTCCAGTTCAAGGGCGAGTGCTACTACACCAAC GGGACGCAGCGCATACGGCTCGTGACCAGATACATCTACAACCGGGAGGAGTACGTGCGC GGGACGCAGCGCATACGGCTCGTGACCAGATACATCTACAACCGGGAGGAGTACGTGCGC 21; Gaps Length 1385; Indels Score 607.8; DB 8; Pred. No. 6.5e-131; 0; Mismatches 57; Query Match
Best Local Similarity 89.6%;
Matches 673; Conservative ø 61 99 105 181 165 241 225 301

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The present sequence was used in the construction of major

600 9 524 584 644 AGACACAACTACGAGGGGCCGGAGACCAGCACCTCCCTGCGGCGCTTGAACAGCCCAAT 464 GTCATGCTGGAGATGACCCCTCATCAGGGAGGTCTACACCTGCCATGTGGAGCATCCC 720 Single chain major histocompatibility complex comprising linked alpha and beta chains - useful for suppressing an immune response to an auto:immune disease, e.g. multiple sclerosis, rheumatoid arthritis, diabetes 345 TACTGGAACAGCCAGCCGGAGATCCTGGAGCGAACGCGGGGCCGAGGTGGACACGCGTGC GTGGCCATCTCCCTGTCCAGGACAGAGCCCTCAACCACACAACACTCTGGTCTGTTCG GTGACAGATTTCTACCCAGCCAAGATCAAAGTGCGCTGGTTCAGGAATGGCCAGGAGGAG GTGACAGATTTCTACCCCAGCCAAGATCAAAGTGCGCTGGTTCAGGAATGGCCAGGAGGAG AGACACAACTACGAGGGCCGGAGACCAGCACCTCCCTGCGGGGGGCGCTTGAACAGCCCAAT GTCGCCATCTCCCTGTCCAGGACAGAGCCCTCAACCACCACACACTCTGGTCTGTTCG ACAGTGGGGGTCTCATCCACACACGCTTATTAGGAATGGGGACTGGACCTTCCAGGTCCTG Construction; major histocompatibility complex; MHC; fusion complex; SCT1 single chain gene; ss. 751 705 AGCCTGAAGAGCCCCATCACTGTGGAGTGGA 735 Wong HC; AGCCTGAAGAGCCCCATCACTGTGGAGTGGA Example 17; Page 137-139; 217pp; English. Location/Qualifiers Rhode PR, Jiao J, Burkhardt M, BP AAT86988 standard; DNA; 1508 97WO-US001617. 96US-00596387 (first entry) SCT1 single chain gene. (DADE-) DADE INT INC. WPI; 1997-402555/37. P-PSDB; AAW29213 WO9728191-A1 30-JAN-1997; 31-JAN-1996; 27-MAR-1998 07-AUG-1997. Synthetic. 405 361 421 465 541 AAT86988; mellitus, 481 525 601 661 721 RESULT 10 Key 셤 셤 g ò ઠે ò g ò ð g ò g ઠ

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complex (sc-MHC) class II complexes that comprise a peptide binding groove, and a modified class II beta 2 chain or covalently linked groove, and a modified class II beta 2 chain or covalently linked cromplexes immunoglobulin (Ig) light chain constant (CI) region. The MHC complexes are useful for detection and analysis of peptide ligands, pathogenic T cells, for functional, cellular and molecular assays. They can be used to identify and isolate T cell receptor and/or MHC agonists and antagonists. They can be used in vivo to compete with pathogenic antigen presenting cells involved in immune-related disorders. They can also be used to arise antibodies and to screen immune cells. It is also use in a method of suppressing an immune response in mammals. The sc-MHC complexes comprising modified class II beta 2 chains and/or 19-CI regions are soluble and provide enhanced yield. These MHC complexes also can contain single antigenic peptides readily isolated from expressing cells in single can compare the particular and analytic complexes also provide a means to detect cells expressing multiple target structures with a single complex. The present sequence represents a DNA encoding a single chain
  61 AGCAGCCCCGGGACTGAGGGCGGAAACTCCATCTGCTTCTCGCCGTCGCTGGAGCACCCG 120
   TACGACAGCGACGTGGGCGAGTACCGCGCGGGGCGAGCTGGGGGCGGCCGAGACGCCGAG 344
   361 TACTGGAACAGCCAGCCGGAGATCCTGGAGCGAACGCGGGCCGAGGTGGACACGGCGTGC 420
  225 GGGACGCAGCGCATACGGCTCGTGACCAGATACATCTACAACCGGGAGGAGTACGTGCGC
  1 ATGGCTCTGCAGATCCCCAGCCTCCTCTCAGCTGCTGTGGTGCTGATGGTGCTG
   121 ATCGTGGTGTCCGGCAGCTGGGACGCTGGGGGCTCACTAGTGCCCCCGAGGCTCTGGA
  105 GCTGCTCACGCTGAAATCAACGAAGCTGGTCGTGCTAGCGGAGGGGGGCGGAAGCGGCGGA
  GGTGGAGGCTCCGAAAGGCATTTCGTGGTCCAGTTCAAGGGCGAGTGCTACTACACCAAC
  GGGGGAAACTCCGAAAGGCATTTCGTGGTCCAGTTCAAGGCGAGTGCTACTACACCAAC
   GGGACGCAGCGCATACGGCTCGTGACCAGATACATCTACAACCGGGAGGAGTACGTGCGC
  invention relates to new single chain major histocompatibility
   chain major histocompatibility complex class I complexes
   Length 1508;
   Sequence 1508 BP; 337 A; 413 C; 441 G; 317 T; 0 U; 0 Other;
immune-related disorder; antigenic peptide; fusion protein;
  Indels
  Wong HC;
  66.2%; Score 607.8; DB 2;
llarity 89.6%; Pred. No. 6.7e-131;
Conservative 0; Mismatches 57;
   ם,
   Jiao
   Burkhardt M,
  IAd/OVA 323-229 MHC fusion protein
   Example 1; Fig 1; 148pp; English.
  98WO-US021520
   (SUNO-) SUNOL MOLECULAR CORP
  PR, Acevedo J,
  WPI; 1999-418411/35.
  Similarity
  P-PSDB; AAY27111
  13-OCT-1998;
  29-OCT-1997;
   06-MAY-1999
  673;
                                  Synthetic.
   Query Match
  181
  165
   241
  301
   Single
   Rhode
 %XGXGXGXGXGXGXXBXABXXBXXBXXBXXBXXBXXBXXBXXBXXBXX
   g
  q
  유
  g
   유
   ò
  셤
  à
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  ð
  ò
  720
   360
  420
   480
  540
   9
  99
   Major histocompatibility complex; MHC; single chain MHC; sc-MHC; Ig; peptide binding groove; immunoglobulin; T cell receptor; immune response;
   AGCAGCCCCGGGACTGAGGGCGGAAACTCCATCTGCTTCTCGCCGTCGCTGGAGCACCCC 120
  121 ATCGTGGTGTCCGGCAGCTGGACGGAGGTGGGGGCTCACTAGTGCCCCGAGGCTCTGGA 180
  300
  284
  344
   404
  464
   524
  584
   644
  704
   65
   GTCATGCTGGAGATGACCCCTCATCAGGGAGAGGTCTACACCTGCCATGTGGAGCATCCC
  1 ATGGCTCTGCAGATCCCCAGCCTCCTCTCAGCTGCTGTGGTGGTGCTGATGGTGCTG
  6 Argecrergeagarececageerecreerereagergergergarecreare
  GGTGGAGGCTCCGAAAGGCATTTCGTGGTCCAGTTCAAGGGCGAGTGCTACTACACCAAC
   GGGGGAAACTCCGAAAGGCATTTCGTGGTCCAGTTCAAGGGCGAGTGCTACTACACCAAC
  GGGACGCAGGCGATACGGCTCGTGACCAGATACATCTACAACCGGGAGGAGTACGTGCGC
  25 GGGACGCAGCGCATACGGCTCGTGACCAGATACATCTACAACCGGGAGGAGTACGTGCGC
   TACGACAGCGACGTGGGCGAGTACCGCGCGGTGACCGAGCTGGGGCGGCCAGACGCCGAG
  TACTGGAACAGCCGGAGATCCTGGAGCGAACGCGGGCCGAGGTGGACACGGCGTGC
   345 TACTGGAACAGCCAGCCGGAGATCCTGGAGCGAACGCGGGCCGAGGTGGACACGCGTGC
   AGACACAACTACGAGGGCCCGGAGACCAGCACCTCCCTGCGGCGCCTTGAACAGCCCAAT
  AGACACAACTACGAGGGCCCGGAGACCAGCACCTCCCTGCGGGGGGCTTGAACAGCCCAAT
  GTCGCCATCTCCCTGTCCAGGACAGAGCCCTCAACCACCACACACTCTGGTCTGTTCG
  GTCGCCATCTCCCTGTCCAGGACAGAGGCCCTCAACCACCACAAAAACACTCTGGTCTGTTCG
   GTGACAGATTTCTACCCAGCCAAGATCAAAGTGCGCTGGTTCAGGAATGGCCAGGAGGAG
   GTGACAGATTTCTACCCAGCCAAGATCAAAGTGCGTTGGTTCAGGAATGGCCAGGAGGAG
  ACAGTGGGGGTCTCATCCACACATTATTAGGAATGGGGACTGGACTTCCAGGTCCTG
  ACAGTGGGGGTCTCATCCACCACAGCTTATTAGGAATGGGGACTGGACCTTCCAGGTCCTG
   GTCATGCTGGAGATGACCCCTCATCAGGGAGGTCTACACACCTGCCATGTGGAGCATCCC
   Gaps
   21;
  Single chain IAd/OVA 323-229 MHC fusion protein encoding DNA.
                                  Sequence 1508 BP; 337 A; 413 C; 441 G; 317 T; 0 U; 0 Other;
   Indels
histocompatibility complex (MHC) fusion complexes
   Score 607.8; DB 2;
Pred. No. 6.7e-131;
0; Mismatches 57;
  751
   AGCCTGAAGAGCCCCATCACTGTGGAGTGGA 735
   AGCCTGAAGAGCCCCATCACTGTGGAGTGGA
  ВР
   Match 66.2%;
Local Similarity 89.6%;
les 673; Conservative C
  AAX89069 standard; DNA; 1508
   (first entry)
  14-SEP-1999
   61
   99
  102
  241
  285
  405
   585
  645
  721
   181
   165
   301
  361
   465
   541
  525
  601
   661
   AAX89069
   421
  481
   705
   Query Match
   Matches
  RESULT 11
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   720
  704
TACTGGAACAGCCAGCCGGAGATCCTGGAGCGAACGCGGGCCGAGCTGGACACGGCGTGC
   AGACACAACTACGAGGGCCGGAGACCAGCCTCCTGCGGGGGCTTGAACAGCCCAAT
   Gredechrerecterechagacadadagecereaacacadaacacacagareregregere
   GTGACAGATTTCTACCCAGCCAAGATCAAAGTGCGCTGGTTCAGGAATGGCCAGGAGGAG
   ACAGTGGGGGTCTCATCCACACACTTATTAGGAATGGGGACTGGACCTTCCAGGTCCTG
  GTCTGCTGGAGATGACCCCTCATCAGGAGAGGTCTACACCTGCCATGTGGAGCATCCC
                      AGACACAACTACGAGGGGCCGGAGACCAGCACCTCCCTGCGGGGGGGCTTGAACAGCCCAAT
   GTGACAGATTTCTACCCAGCCAAGATCAAAGTGCGCTGGTTCAGGAATGGCCAGGAGGAG
   ACAGTGGGGGTCTCATCCACACATTATTAGGAATGGGGACTGGACCTTCCAGGTCCTG
   GTCATGCTGGAGATGACCCCTCATCAGGAGGGTCTACACCTGCCATGTGGAGCATCCC
  MHC; major histocompatibility complex; gene therapy; fusion complex; peptide-binding groove; T cell modulation; class II MHC; vaccine; autoimmune disorder; multiple sclerosis; rheumatoid arthritis; insulin-dependent diabetes mellitus; myasthenia gravis; immunogen; chronic allergy; mouse; ds; I-Ad; gene.
  ÅĊ;
   Edwards
   Mouse MHC I-Ad/Ova 323-339 synthetic gene SCT1.
  751
  Grammer S,
  AGCCTGAAGAGCCCCATCACTGTGGAGTGGA
  Weidanz JA,
  ACA60743 standard; DNA; 1508 BP
   94US-00283302.
95US-00382454.
97US-00776084.
  06-JUL-2001; 2001US-00900379
   entry)
  Jiao JJJ;
  (DADE-) DADE INT INC.
   (first
  2003-341126/32.
  Rhode PR,
  US2002198144-A1
  Chavaillaz P,
   01-FEB-1995;
17-JAN-1997;
  16-JUN-2003
   26-DEC-2002
  Synthetic.
   721
  405
   585
                      421
  481
   465
   541
  525
   601
   199
  645
  705
  ACA60743;
  НĊ,
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complex (I) comprising an MHC molecule that contains a peptide-binding CC groove, and a presenting peptide covalently (e.g. an antigenic peptide) linked to the MHC molecule, where (I) is capable of modulating the correct of a T cell. Also included are a DNA construct coding for the activity of a T cell. Also included are a DNA construct coding for the complex, where the MHC molecule is a class IN MHC (e.g. mounes I-Ad or I-As, or human HLA-DRI (human laukocyce antigen-DRI)), a multivalent MHC fusion complex comprising two or more linked complexes, identifying a cut to the cells cloning vectors that each contain the fusion complex and selecting host cells that cam modulate the activity of T cells (involving introducing into host cells under conditions suitable for expression of the MHC fusion complex, and selecting host cells that expression of the CMHC fusion complex, and selecting host cells that capas MHC fusion complex that codes for the alpha and beta chains of the MHC fusion complex that codes for a T cell costimulatory factor and beta chains of the MHC fusion complex. The DNA constructs can contain the terecologous leader peptide sequences and Kozak sequence for efficient cresponse in a mammal (including vaccinating a mammal against a targeted disorder, by administering DNA sequence comprising a mammal against a targeted disorder, by administering an expression vector, encoding a till leaph MHC molecule that contains a transmembrane domain, and complex which is a single chain fusion complex which is a single chain fusion complex which is a single chain fusion complex which is a single chain fusion complex which is a contain or because and suppressing an immune response in a mammal (including accinating a peptide that is a T cell receptor (TCR) antagonist or partial coding for the fusion complex which is a single chain disorder the methods are useful for identifying a peptide that can modulate the contain of a mammal gainst a targeted disorders multiple and an ammal. The disorders include an autoimmune com
  180
  61 AGCAGCCCCGGGACTGAGGCCGGAAACTCCATCTGCTTCTCGCCGTCGCTGGAGCACCCG 120
   104
  164
  300
  360
   420
   480
  224
   284
   344
  404
   65
   TACGACAGCGACGTGGGCGGGTGACCGCGGGGTGACCGAGCTGGGGCGGCCAGACGCGGCGGCGGG
  1 ATGGCTCTGCAGATCCCCAGCCTCCTCTCAGCTGCTGGTGGTGGTGCTGATGGTGCTG
   121 ATCGTGGTGTCCGGCAGCTGGGACGGGGGGGGCTCACTAGTGCCCCGAGGCTCTGGA
  GCTGCTCACGCTGAAATCAACGAAGCTGGTCGTGCTAGCGGAGGGGGGCGGAAGCGGCGGA
  GGGGGAAACTCCGAAAGGCATTTCGTGGTCCAGTTCAAGGGCGAGTGCTACTACACCAAC
  GGGACGCAGCGCATACGGCTCGTGACCAGATACATCTACAACCGGGAGGAGTACGTGCGC
  TACTGGAACAGCCAGCCGGAGATCCTGGAGCGAACGCGGGCCCGAGGTGGACACGGCGTGC
   GGTGGAGGCTCCGAAAGGCATTTCGTGGTCCAGTTCAAGGGCGAGTGCTACTACACCAAC
   GGGACGCAGCGCATACGGCTCGTGACCAGATACATCTACAACCGGGAGGAGTACGTGCGC
  TACGACAGCGACGTGGGCGAGTACCGCGCGGTGACCGAGGTGGGGCGGCCGAGACGCCGAG
   AGACACAACTACGAGGGCCGGAGACCAGCACCTCCCTGCGGGGGCGTTGAACAGCCCAAT
   TACTGGAACAGCCAGCCGGAGATCCTGGAGCGAACGCGGGCCGAGGTGGACACGGCGTGC
   Gaps
  21;
  Sequence 1508 BP; 337 A; 413 C; 441 G; 317 T; 0 U; 0 Other;
  Indels
  57;
   Score 607.8; DB 8;
Pred. No. 6.7e-131;
  0; Mismatches
   66.2%;
89.6%;
  Matches 673; Conservative
   Similarity
  invention
  105
   181
  165
  241
   225
   285
  345
   Query Match
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   361
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  Local
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The invention relates to a major histocompatibility complex (MHC) fusion

Example 17; Fig 28; 126pp; English.

allergies

Novel major histocompatibility complex fusion complex having presenting peptide covalently linked to MHC molecule containing peptide-binding groove, used for suppressing immune response in multiple sclerosis,

P-PSDB; ABU72107

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61 AGCAGCCCCGGGACTGAGGGCGGAAACTCCATCTGCTTCTCGCCGTCGCTGGAGCACCCG 120
  105 gcráctcacectéaaatcaaceaaecréercerectaeceaaeeeeeeaeeceeea 164
  6 ATGGCTCTGCAGATCCCCAGCCTCCTCTCAGCTGCTGTGGTGGTGCTGATGGTGCTG
  1 ATGGCTCTGCAGATCCCCAGCCTCCTCTCAGCTGCTGTGGTGGTGGTGGTGGTGCTG
   121 ATCGTGGTGTCCGGCAGCTGGACGGAGGTGGGGGCTCACTAGTGCCCCGAGGCTCTGGA
  Matches 672; Conservative
  Local Similarity
  241
   301
  181
  Query Match
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   GTCATGCTGGAGATGACCCCTCATCAGGGAGAGGTCTACACCTGCCATGTGGAGCATCCC 720
  644
   Location/Qualifiers
6. .1382
6. .1382
6. .86
7.tag= b
7.tag= b
7.tag= b
7.tag= In 1-4d beta chain leader
7.tag= In 1-4d beta chain leader
  MHC; major histocompatibility complex; PCR; polymerase chain reaction; T cell activity modulator; antagonist; immune disorder; allergy; multiple sclerosis; insulin-dependent diabetes mellitus; rheumatoid arthritis; myasthenia gravis; ds.
                  465 GTCGCCATCTCCCTGTCCAGGACAGAGGCCCTCAACCACCACAACACTCTGGTCTGTTCG
  CAGGAATGGCCAGGAGGAG
  525 GIGACAGAITICIACCCAGCCAAGAICAAGIGCGCIGGIICAGGAAIGGCCAGGAGGG
   ACAGTGGGGGTCTCATCCACACAGCTTATTAGGAATGGGGACTGGACCTTCCAGGTCCTG
   585 ACAGTGGGGGTCTCATCCACACAGCTTATTAGGAATGGGGACTGGACCTTCCAGGTCCTG
  /label= I-Ad alpha2
/note= "murine MHC class II I-Ad gene alpha-2 domain"
  gene alpha-2 domain"
   I-Ad gene beta-1 domain"
  domain"
  Vector SCE1-derived single chain gene encoding MHC fusion complex.
   gene beta-2
   /label= OVA_323-339
/note= "chicken ovalbumin residues 323-339"
138. .167
/note= "10 residue linker peptide"
  rtag= g
hote= "24 residue peptide linker"
007. .1067
  GTGACAGATTTCTACCCCAGCCAAGATCAAAGTGCGCTGGTT
  label= I-Ad beta2
note= "murine MHC class II I-Ad (35. .806
   *tag= h
|label= I-Ad alphal
|note="murine MHC class II I-Ad |
|068. 1352
   AGCCTGAAGAGCCCCATCACTGTGGAGTGGA 751
   Η
  '*tag= e
label= I-Ad_betal
'note= "murine MHC class
153. .734
   BP
   AAT17588 standard; DNA; 1382
  (first entry)
   .452
  equence"
  /*tag=
/label=
  *tag=
   misc_feature
   misc_feature
  misc_feature
  misc_feature
   misc_feature
   misc_feature
  misc_feature
  26-SEP-1996
   sig_peptide
  Synthetic
   721
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   AAT17588;
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180

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Gaps

21;

Indels

66.0%; Score 606.2; DB 2; 89.5%; Pred. No. 1.5e-130; iive 0; Mismatches 58;

Length 1382;

GGTGGAGGCTCCGAAAGGCATTTCGTGGTCCAGTTCAAGGGCGAGTGCTACTACACCAAC 240

GGGACGCAGCGCATACGGCTCGTGACCAGATACATCTACAACCGGGAGGAGTACGTGCGC 300

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AAT17588 encodes a murine MHC fusion complex capable of modulating T cell activity encoded by the vector SCEI. The MHC fusion complex comprises at least one MHC molecule containing a peptide-binding groove and a presenting peptide covalently linked to the MHC molecule and opt. a crassing peptide covalently linked to the MHC molecule and opt. a crassing and transmembrane domain. DNA encoding a MHC fusion complex may be cloned into a host cell to express the complex. The transformed cells may then cativity. DNA encoding a MHC fusion complex or a single chain fusion activity. DNA encoding a MHC fusion complex or a single chain fusion complexes may be used to suppress an immune response in an enimal suffering from an immune disorder e.g. multiple sclerosis, insulin dependent diabetes mellitus, rheumatcoid arthritis, myasthenia gravis or chronic allergies. The complexes may also be used in the treatment of livestock and pets such as casts and dogs. The MHC fusion complexes can be produced such that they contain a single antigenic peptide including one of known structure, additionally a wide range of peptides can be contain a single antigenic peptide including one contain a single antigenic peptide including one contain a single and complexes can be produced for T cell interaction
  Major histocompatability complex fusion complex for modulating T cel activity - used in the treatment of immune disorders, e.g. multiple sclerosis, IDDM and rheumatoid arthritis.
  Sequence 1382 BP; 320 A; 374 C; 404 G; 284 T; 0 U; 0 Other;
   Edwards AC;
   Grammer S,
   Example 17; Fig 29; 210pp; English
   Weidanz JA,
  /*tag= j
/note= "EE tag"
  95WO-US009816
   94US-00283302
95US-00382454
.1379
  (DADE-) DADE INT INC.
   Wong HC, Rhode PR,
Chavaillaz P, Jiao
   WPI; 1996-129343/13.
P-PSDB; AAR98907.
    misc_feature
  31-JUL-1995;
  29-JUL-1994;
  WO9604314-A1
  01-FEB-1995;
   15-FEB-1996.
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                 420
                                404
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   540
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  999
  720
   524
   584
   644
TACGACAGCGACGTGGCGAGTACCGCGCGGTGACCGAGCTCGGGCGGCGACCACACGCCGAG 344
   645 GTCATGCTGGAGATGACCCCTCATCAGGGAGAGGTCTACACCTGCCATGTGGAGCATCCC 704
   /label= I-Ad_beta_chain_leader
/note= "murine MHC class II I-Ad gene beta chain leader
sequence"
87. .137
  MHC; major histocompatibility complex; PCR; polymerase chain reaction; T cell activity modulator; antagonist; immune disorder; allergy; multiple sclerosis; insulin-dependent diabetes mellitus; rheumatoid arthritis; myasthenia gravis; ds.
                        AGACACACTACGAGGGCCCGCAGACCAGCACCTCCCTGCGGCGCCTTGAACAGCCCAAT
   GTCGCCATCTCCCTGTCCAGGACAGAGGCCTCAACCACACAACACTCTGGTCTGTTCG
  GTGACAGATTTCTACCCAGCCAAGATCAAAGTGCGCTGGTTCAGGAATGGCCAGGAGGAG
   525 GTGACAGATTICTACCCAGCCAAGATCAAAGTGCGCTGGTTCAGGAATGGCCAGGAGAG
  ACAGTGGGGGTCTCCACCACACACTTATTAGGAATGGGGACTGGACCTTCCAGGTCCTG
  GTCATGCTGGAGATGACCCCTCATCAGGAGGTCTACACCTGCCATGTGGAGCATCCC
                TACTGGAACAGCCAGCCGGAGATCCTGGAGCGAACGCGGGCCCGAGGTGGACACGGCGTGC
  /note= "murine MHC class II I-Ad gene beta-2 domain"
735. .806
   domain"
  Vector SSC1-derived single chain gene encoding MHC fusion complex
  beta-1
   /*tag= c
/label= OVA 323-339
/note= "chicken ovalbumin residues 323-339"
138. .167
/*tag= d
/note= "10 residue linker peptide"
  gene
   /label= I-Ad betal
/note= "murine MHC class II I-Ad
453. .734
  705 AGCCTGAAGAGCCCCATCACTGTGGAGTGGA 735
   721 AGCCTGAAGAGCCCCATCACTGTGGAGTGGA 751
   location/Qualifiers
  '*tag= f
'label= I-Ad_beta2
   AAT17586 standard; DNA; 1385 BP
   (first entry)
  ៧
   168. .452
  *tag= e
  . .1385
  /*tag=
6.86
/*tag=
  /*tag=
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  misc_feature
   misc feature
   misc_feature
   misc_feature
  26-SEP-1996
  sig_peptide
                361
  421
  465
   Synthetic
  541
  481
  AAT17586;
  601
  661
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AAT17586 encodes a murine MHC fusion complex capable of modulating T cell activity encoded by the vector SSC1. The MHC fusion complex comprises at least one MHC molecule containing a peptide-binding groove and a presenting peptide covalently linked to the MHC molecule and opt. a transmembrane domain. DNA encoding a MHC fusion complex may be cloned into a host cell to express the complex. The transformed cells may then ce used to identify peptides that modulate, pref. antegonise, T cell activity. DNA encoding a MHC fusion complex or a single chain fusion molecule may be used to vaccinate a mammal against a targeted disorder. The fusion complexes may be used to suppress an immune response in an animal suffering from an immune disorder e.g. multiple sclerosis, insulin dependent diabetes mellitus, rheumatoid arthritis, myasthenia gravis or chronic allergies. The complexes may also be used in the treatment of livestock and pets such as cards and dogs. The MHC fusion complexes can be produced such that they contain a single antigenic peptide including one of known structure, additionally a wide range of peptides can be presented for T cell interaction
   61 AGCAGCCCCGGGACTGAGGGCGGAAACTCCATCTGCTTCTCGCCGTCGCTGGAGCACCCG 120
   181 GGTGGAGGCTCCGAAAGGCATTTCGTGGTCCAGTTCAAGGGCGAGTGCTACTACACAAC 240
  65
   ATGCTCTGCAGATCCCCAGCCTCCTCTCTCAGCTGCTGTGGTGGTGGTGATGGTGTG
  ------criaagtatcrcrcaggctgttcac
  gerácreacecránantenación de respectados de desegencia de conferencia  1 ATGCCTCTGCAGATCCCCAGCCTCCTCTCAGCTGCTGTGGTGGTGCTGATGGTGCTG
   121 ATCGTGGTGTCCGGCAGCTGGACGGGGGGGCCTCACTAGTGCCCCCGAGGCTCTGGA
  gene alpha-2 domain"
  II I-Ad gene alpha-2 domain"
   Gaps
   Major histocompatability complex fusion complex for modulating T ce. activity - used in the treatment of immune disorders, e.g. multiple sclerosis, IDDM and rheumatoid arthritis.
   21;
  Score 606.2; DB 2; Length 1385; Pred. No. 1.5e-130;
   Sequence 1385 BP; 316 A; 384 C; 398 G; 287 T; 0 U; 0 Other;
  Edwards AC;
   Indels
note= "24 residue peptide linker"
  II I-Ad
   58;
   Grammer S,
   /*tag= h
/label= I-Ad alpha1
/label= murine MHC class II
1068. 1352
/*tag= i.
/label= I-Ad_alpha2
/lote= murine MHC class III
   0; Mismatches
   Example 17; Fig 27; 210pp; English
  /*tag= j
/note= "6xHis tag"
  Rhode PR, Weidanz JA,
  66 AGCAGCCCAAGGAC----
   95WO-US009816
   94US-00283302
95US-00382454
  Query Match
Best Local Similarity 89.5%;
Matches 672; Conservative (
  .1067
  Jiao J;
   (DADE-) DADE INT INC
   WPI; 1996-129343/13.
  P-PSDB; AAR98905.
  Chavaillaz P,
  31-JUL-1995;
  misc_feature
   misc_feature
   misc_feature
   29-JUL-1994;
   01-FEB-1995;
   WO9604314-A1
  15-FEB-1996
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1 ATGGCTCTGCAGATCCCCAGCCTCCTCTCAGCTGCTGTGGTGGTGCTGATGGTGCTG
  31-JUL-1995;
  29-JUL-1994;
   01-FEB-1995;
  misc_feature
  misc_feature
  misc_feature
   misc_feature
   misc feature
  WO9604314-A1
   HC,
   Wong
   ò
   480
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  9
  360
  420
  720
  284
  404
   464
   524
  584
  644
  /*tag= a
6. .86
/*tag= b
/label= I-Ad beta chain leader
/note= "murine MHC class II I-Ad gene beta chain leader
  MHC; major histocompatibility complex; PCR; polymerase chain reaction; T cell activity modulator; antagonist; immune disorder; allergy; multiple sclerosis; insulin-dependent diabetes mellitus; rheumatoid arthritis; myasthenia gravis; ds.
225 GGGACGCAGCGCATACGGCTCGTGACCAGATACATCTACAACCGGGAGGAGTACGTGGCC
  AGACACAACTACGAGGGGCCGGAGACCAGCACCTCCCTGCGGCGCGCTTGAACAGCCCAAT
  GGGACGCAGCGCATACGGCTCGTGACCAGATACATCTACAACCGGGAGGAGTACGTGCGC
  TACGACAGCGACGTGGGCGAGTACCGCGCGGGGGCGGCGGCGGCCGAGACGCCGAG
  TACTEGAACAGCCAGCCGGAGATCCTGGAGCGAACGCGGGCCCGAGGTGGACACGGCGTGC
  AGACACAACTACGAGGGCCCGGAGACCAGCACCTCCCTGCGGCGGCGCTTGAACAGCCCAAT
   gresecearerecersteassacasassaceretaaceaceacaacaereresereres
   GTGACAGATTTCTACCCAGCCAAGATCAAAGTGCGCTGGTTCAGGAATGGCCCAGGAGGAG
  GTGACAGATTTCTACCCAGCCAAGATCAAAGTGCGCTGGTTCAGGAATGGCCAGGAGGAG
  ACAGTGGGGGTCTCCACCACACCTTATTAGGAATGGGGACTGGACCTTCCAGGTCCTG
  acagregesercrearceacacerrarraggaaresesacresacerrecasercers
  GTCATGCTGGAGATGACCCCTCATCAGGGAGAGGTCTACACCTGCCATGTGGAGCATCCC
  Vector SCT1-derived single chain gene encoding MHC fusion complex.
  /*tag= c
/label= OVA 323-339
/note= "chicken ovalbumin residues 323-339"
138. .167
  /*tag= d
/note= "10 residue linker peptide"
168. .452
  751
  AGCCTGAAGAGCCCCATCACTGTGGAGTGGA 735
  AGCCTGAAGAGCCCCATCACTGTGGAGTGGA
   Location/Qualifiers
6. .1508
   AAT17587 standard; DNA; 1508 BP
  (first entry)
   sequence"
   87. .137
   misc_feature
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  misc_feature
   26-SEP-1996
   sig_peptide
  Synthetic
  345
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                       165
  301
  285
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   421
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AAT17587 encodes a murine MHC fusion complex capable of modulating T cell activity encoded by the vector SCT1. The MHC fusion complex comprises at least one MHC molecule containing a peptide-binding groove and a presenting peptide covalently linked to the MHC molecule and opt. a cransmembrane domain. DNA encoding a MHC fusion complex may be cloned into a host cell to express the complex. The transformed cells may then be used to identify peptides that modulate, pref. antegoniae, T cell activity. DNA encoding a MHC fusion complex or a single chain fusion complexes may be used to suppress an immune response in an entire suffering from an immune disorder e.g. multiple sclerosis, insulince animal suffering from an immune disorder e.g. multiple sclerosis, insulince chronic allergies. The complexes may also be used in the treatment of chronic allergies. The complexes may also be used in the treatment of chronic allergies. The complexes may also be used in the treatment of chronic allergies. The complexes may also be used in the treatment of chronic allergies. The complexes may also be used in the treatment of chronic allergies. The state and dogs. The MHC fusion complexes can be conduced such that they contain a single antigenic peptide including one of known structure, additionally a wide range of peptides can be complexed.
   ä
  /*tag= j
/label= I-Ad_alpha-TM
/note= "murine MHC class II I-Ad gene alpha-transmembrane
   Major histocompatability complex fusion complex for modulating T cell activity - used in the treatment of immune disorders, e.g. multiple sclerosis, IDDM and rheumatoid arthritis.
   /*teg= 1
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1353. 1505
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/note= "murine MHC class II I-Ad gene alpha-2 domain"
   Gaps
/*tag= e
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453. .734
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/label= I-Ad_beta2
/note= "murine MHC class II I-Ad gene beta-2 domain"
735. 806
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29-SEP-1999

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linear

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Rhode, P. R., Jao, J. -A., Burkhardt, M.

MHC molecules and uses thereof

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70.7%
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LOCUS
DEFINITION
ACCESSION
VERSION
KEYWORDS
SOURCE
   ORGANISM
   AUTHORS
TITLE
JOURNAL
FEATURES
  RESULT 15
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Aavi2068 Murine IA
Aat104269 Hybrid IA
Aat17588 Vector SC
Aat86989 SCE1 sing
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Abj99031 MBP 90-10
  Abk63510 Rat seque
Adb57995 Toxicity
Abt41775 Toxicity
   Recombinant protein; alpha chain; beta chain; MHC; immunoglobulin; major histocompatibility complex; Fc region; antigen; I lymphocyte; immunostimulant; vaccine; infection; tumour; ss.
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-OUTFWT=ptc -NORM=ext -HEAPSIZE=500 -MINLEN=0 -MAXLEN=2000000000 -HOST=abss05p
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-WARN TIMEOUT=30 -THREADS=1 -XGAPOP=10 -XGAPEXT=0, 5 -FGAPEXT=7
   Aaf55099 DNA encod
Aat04262 Hybrid IA
Adg31225 I-Ab(beta
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  Pred. No. is the number of results predicted by chance to have a score greater than or equal to the score of the result being printed, and is derived by analysis of the total score distribution.
  1620
1 MALQIPSLLLSAAVVVLMVL.....LKWKLQALKKKLAQHHHHHH 306
   Description
  10489840
               GenCore version 5.1.9
Copyright (c) 1993 - 2006 Biocceleration Ltd.
   OM protein - nucleic search, using frame_plus_p2n model
  hits satisfying chosen parameters:
   5244920 segs, 3486124231 residues
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   Post-processing: Minimum Match 0%
Maximum Match 100%
Listing first 45 summaries
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AAT04262
ADQ31225
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Xgapop 10.0, Xgapext
Ygapop 10.0, Ygapext
Fgapop 6.0, Fgapext
Delop 6.0, Delext
  geneseqn2003ds:*geneseqn2004as:*geneseqn2004bs:*geneseqn2005s:*geneseqn2006s:*
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  US-10-048-116B-6
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  921
893
945
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   100.0
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Database :

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   GTGACAGATTTCTACCCAGCCAAGATCAAAGTGCGCTGGTTCAGGAATGGCCAGGAGGG
   GlyGlyGlySerThrThrAlaProSerAlaGlnLeuLysLysLysLysLeuGlnAlaLeu
   GTCATGCTGGAGATGACCCCTCATCAGGGAGGGTCTACACCCTGCCATGTGGAGCATCCC
   SerLeuLysSerProlleThrValGluTrpArgAlaGlnSerGluSerAlaArgSerLys
   ValThrAspPheTyrProAlaLysIleLysValArgTrpPheArgAsnGlyGluGlu
   Thr ValGly ValSerSerThrGlnLeuIleArgAsnGlyAspTrpThrPheGlnValLeu
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  te primer binding site (primer #233)". .74)
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complement(119, 172)
   /*tag= b
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  Location/Qualifiers
  1. .16
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  CATCATCATCATCAT 918
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  ВP
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   Hybrid IA beta chain gene.
   .828
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  661
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  721
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  The specification describes soluble recombinant proteins that comprise at least a dimer formed from the alpha and beta-chains of MHC (major histocompatibility complex) Class I and II molecules in which at least one chain has, attached to its C-terminus, at least part of the Fc region of comparinc peptide, are used to count and/or purify antigen-reactive T clymphocytes and to characterize their phenotype, e.g. in preclinical evaluation of vaccines. They are also used as immunostimulants, count and determine phenotype of autoreactive T cells in subjects with, count and determine phenotype of autoreactive T cells in subjects with, count and determine phenotype of autoreactive T cells in subjects with, cor at risk of developing, and (to purify and/or enrich Ag-reactive T cells from cell cultures or patient samples, for use in subsequent curative or preventative cellular therapy. The present sequence encodes a recombinant protein of the invention, comprising a beta chain of MHC molecules
  140
   180
   240
  GlyThrGlnArglleArgLeuValThrArgTyrlleTyrAsnArgGluGluTyrValArg 100
  300
   120
   360
   ArgHisAsnTyrGluGlyProGluThrSerThrSerLeuArgArgLeuGluGlnProAsn 160
   20
  9
   40
  9
   80
  GlyGlyGlySerGluArgHisPheValValGlnPheLysGlyGluCysTyrTyrThrAsn
   TyrAspSerAspValGlyGluTyrArgAlaValThrGluLeuGlyArgProAspAlaGlu
  TyrTrpAsnSerGlnProGluIleLeuGluArgThrArgAlaGluValAspThrAlaCys
  MetAlaLeuGlnIleProSerLeuLeuLeuSerAlaAlaValValValLeuMetValLeu
   SerSerProGlyThrGluGlyGlyAsnSerIleCysPheSerProSerLeuGluHisPro
   GGTGGAGGCTCCGAAAGGCATTTCGTGGTCCAGTTCAAGGGCGAGTGCTACTACACCAAC
  GGGACGCAGCGCATACGCCTCAGCAGATACATCTACAACCGGGAGGAGTACGTGCGC
   S
   New soluble recombinant protein, useful e.g. as immunostimulant, comprises dimeric major histocompatibility complex molecule fused immunoglobulin Fc region.
  Sequence 921 BP; 214 A; 265 C; 286 G; 156 T; 0 U; 0 Other;
   921
306
0
0
0
   Length:
Matches:
Conservative:
Mismatches:
Indels:
Gaps:
   US-10-048-116B-6 (1-306) x AAF55099 (1-921)
  Example 1; Page 34-35; 43pp; French.
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   6.68e-144
1620.00
100.0%
100.0%
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Best Local Similarity:
   Glaichenhaus N,
   Alignment Scores:
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   61
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   61
  81
  241
  121
   101
  Query Match:
DB:
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405 140 465 160 525

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I-Ab(beta)-Cholera toxin B subunit-leucine zipper (LZ)-BirA fusion cDNA
  class II major histocompatibility complex; MHC; CD4+ T-cell detection; flow cytometry; mucous membrane invasive antigen; I-Ab(beta)-cholera toxin B subunit-leucine zipper-BirA fusion; CTB; ss;
  GGTGGAGGCTCCGAAAGGCATTTCGTGGTCCAGTTCAAGGGCGAGTGCTACTACACCAAC
   TyrAspSerAspValGlyGluTyrArgAlaValThrGluLeuGlyArgProAspAlaGlu
   ThrValGlyValSerSerThrGlnLeuIleArgAsnGlyAspTrpThrPheGlnValLeu
   ValMetLeuGluMetThrProHisGlnGlyGluValTyrThrCysHisValGluHisPro
     41 IleValValSerGlySerTrpAspGlyGlyGlyGlySerLeuValProArgGlySerGly
                            ---GAGATCAATGAGGCTGGCAGAGGAGGTGGGGGCTCACTAGTGCCCCGAGGCTCTGGA
   GlyGlyGlySerGluArgHisPheValValGlnPheLysGlyGluCysTyrThrAsn
  GlyThrGlnArgIleArgLeuValThrArgTyrIleTyrAsnArgGluGluTyrValArg
  GGGACGCAGCGCATACGCCTCGTGACCAGATACATCTACAACCGGGAGGAGTACGTGCGC
  TyrTrpAsnSerGlnProGluIleLeuGluArgThrArgAlaGluValAspThrAlaCys
   TACTGGAACAGCCAGCCGGAGATCCTGGAGCGAACGCGGGCCCAAGGTGGACACGCCGTGC
   ArgHisAsnTyrGluGlyProGluThrSerThrSerLeuArgArgLeuGluGlnProAsn
   AGACACAACTACGAGGGCCGGAGACCAGCACCTCCCTGCGGCGCGCTTGAACAGCCCAAT
  ValAlaIleSerLeuSerArgThrGluAlaLeuAsnHisHisAsnThrLeuValCysSer
  ValThrAspPheTyrProAlaLysIleLysValArgTrpPheArgAsnGlyGlnGluGlu
  GTGACAGATTTCTACCCAGCCAAGATCAAAGTGCGCTGGTTCAGGAATGGCCAGGAGGAG
  ACAGTGGGGGTCTCATCCACACCTATTAGGAATGGGGACTGGACCTTCCAGGTCCTG
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leucine zipper (LZ)-BirA fusion cDNA"
  Location/Qualifiers
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   (first entry)
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Unidentified.
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   07-OCT-2004
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   ADQ31225;
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   This sequence represents a hybrid IA beta chain gene, containing the chicken ovalbumin peptide (cOVA). This sequence was used in the construction of a hybrid IA alpha beta dimer. The encoded protein (pIAd-OVA) was found to be morre stable than the IA alpha beta dimer. The stability was decreased by the addition of a MHC groove specific binding peptide (e.g. see ARR82527), ARR82528 and ARR82511, compared to an increase seen on the addition of a MHC binding peptide to IE k/d-MCC. These complexes may be used to regulate an immune response. The complexes are capable of being recognised by a TCR alone or in combination with purposes and experimental purposes. They can also be used as reagents for the treatment of diseases including autoimmune disease, immunodeficiency
   168
  Peptide-MHC complex comprising antigenic peptide, linker and MHC segment - useful as reagents for the treatment of diseases including auto-immune diseases, immuno-stimulatory diseases or graft-host rejection.
   40
   SerSerProGlyThrGluGlyGlyAsnSerIleCysPheSerProSerLeuGluHisPro
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  note= "probable primer binding site (primer #232)
   diseases, immunoproliferation diseases, and graft-host rejection
  Sequence 893 BP; 204 A; 239 C; 275 G; 175 T; 0 U; 0 Other;
  893
6 42
7 7
1
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Matches:
Conservative:
Mismatches:
Indels:
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   21
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DB:
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240 765 260

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180

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600
   9
  720
   259
   299
  Peptide-Class II major histocompatibility complex (MHC) composite, useful for detecting antigen specific CD4+ T-cell, comprises antigen peptide containing epitope of mucous membrane invasive protein, and extracellular
   834
  CTTAAGAAAAAGAACGCTCAGCTGAAGTGGAAACTTCAAGCCCTCAAGAAGAAACTCGCC 894
  /*tag= a
/product= "I-Ab(beta)-Escherichia coli heat-labile toxin
B subunit (LTB)-leucine zipper (LZ)-BirA fusion protein"
  class II major histocompatibility complex; MHC; CD4+ T-cell detection; flow cytometry; mucous membrane invasive antigen; I-Ab(beta)-heat-labile toxin B subunit-leucine zipper-BirA fusion; LTB.
  GluThrValGlyValSerSerThrGlnLeuIleArgAsnGlyAspTrpThrPheGlnVal
   ProSerLeuLysSerPro11eThrValGluTrpArgAlaGlnSerGluSerAlaArgSer
     AsnvalAlaileSerLeuSerArgThrGluAlaLeuAsnHisHisAsnThrLeuValCys
  TCAGTGACAGATTCTACCCAGCCAAGATCAAAGTGCGCTGGTTCCGGAATGGCCAGGAG
   GAGACGCTCGCGCGCTCTTCCACCCAGCTTATTAGGAATGGGGACTGGACCTTCCAGGTC
   LeuValMetLeuGluMetThrProHisGlnGlyGluValTyrThrCysHisValGluHis
  creditariecredadareacecereecesesasasasereracereresesesear
   LeuLysLysAsnAlaGlnLeuLysTrpLysLeuGlnAlaLeuLysLysLysLeuAla
                                    AATGTCGTCATCTCCTGTCCAGGACAGAGGCCCTCAACCACCACAACACTCTGGTCTGC
   SerValThrAspPheTyrProAlaLyslleLysValArgTrpPheArgAsnGlyGlnGlu
   721 CCCAGCCTGAAGAGCCCCATCACTGTGGAGTGGAGGGCACAGTCGTCAGCA-----GAC
   LysGlyGlyGlyGlySerThrThrAlaProSerAlaGlnLeuLysLysLysLeuGlnAla
   I-Ab(beta)-E. coli heat-labile toxin B subunit-LZ-BirA fusion cDNA
  ठ
  GIJUTSU INCUBATION CENT
   Location/Qualifiers
   ₽.
   ADQ31228 standard; cDNA; 915
  03-DEC-2003; 2003JP-00404367.
   03-DEC-2002; 2002JP-00351818
  07-OCT-2004 (first entry)
   GlnHisHisHis 303
  CAGCTGCATCAT 906
  (SENT-) SENTAN KAGAKU
  WPI; 2004-546819/53.
P-PSDB; ADQ31227.
  coli.
   JP2004196789-A.
  Escherichia
Unidentified
  15-JUL-2004
  ADQ31228;
                                      481
   541
  200
   601
   240
   775
   160
   180
   220
  661
  260
   280
  835
   300
  895
   gene
   RESULT
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   The invention relates to a novel class II major histocompatibility complex (MHC) antigenic peptide composite comprising a peptide containing the T-cell antigenic determinant of a mucous membrane invasive protein and the extracellular region of a class II MHC molecule or at least part of the extracellular region of the class II MHC molecule having an amino acid sequence comprising one or more deletions, substitutions or additions. The molecule of the invention may be useful for detecting an antigen-specific CD4+ T-cell by flow cytometry and for presenting a microorganism-derived mucous membrane invasive protein as an antigen. The method of the invention enables efficient detection of antigen-specific activation of CD4+ T-cells in the mucous membrane. The current sequence is that of the class II major histocompatibility complex-related I-
  Peptide-Class II major histocompatibility complex (MHC) composite, useful for detecting antigen specific CD4+ T-cell, comprises antigen peptide containing epitope of mucous membrane invasive protein, and extracellular
  119
   360
  480
   180
   240
  300
  CysArgHisAsnTyrGluGlyProGluThrSerThrSerLeuArgArgLeuGluGlnPro 159
  123
   19
  59
  79
   66
  63
   LeuSerSerProGlyThrGluGlyGlyAsnSerIleCysPheSerProSerLeuGluHis 39
  ::: |||::::||||
grgretcrgaagctccctggaggttcctacatgccaaagctgacaggacactgatggtg
   ProlleValValSerGlySerTrpAspGlyGlyGlyGlyGerLeuValProArgGlySer
  ArgTyrAspSerAspValGlyGluTyrArgAlaValThrGluLeuGlyArgBroAspAla
   GAGTACTGGAACAGCCGGGGGGGGAGATCCTGGAGCGAACGGGGCCGAGGCTGGACACGGTG
  MetAlaLeuGlnIlePro---SerLeuLeuLeuSerAlaAlaValValValLeuMetVal
   GCGATCGCGGCCATCAGCATGGCGAACGGAGGTGGTGGCTCC---GGTGGAGGGGGAAGT
  GlyGlyGlyGlySerGluArgHisPheValValGlnPheLysGlyGluCysTyrTyrThr
   AsnGlyThrGlnArglleArgLeuValThrArgTyrIleTyrAsnArgGluGluTyrVal
  AACGGGACGCAGCGATACGATATGTGACCAGATACATCTACAACCGGGAGGAGTACGTG
  GluTyrTrpAsnSerGlnProGluIleLeuGluArgThrArgAlaGluValAspThrAla
   U; 0 Other;
   9 2 4 4 4 6
2 4 4 4 5
3 4
                                (SENT-) SENTAN KAGAKU GIJUTSU INCUBATION CENT KK
   Length:
Matches:
Conservative:
Mismatches:
Indels:
Gaps:
   G; 165 T; 0
  Example 1; SEQ ID NO 10; 30pp; Japanese.
   US-10-048-116B-6 (1-306) x ADQ31225 (1-945)
   294
   BP; 230 A; 256 C;
   2.27e-107
1235.00
84.9%
/: 80.3%
03-DEC-2002; 2002JP-00351818
   CDNA of the invention
   WPI; 2004-546819/53.
P-PSDB; ADQ31224.
   Percent Similarity:
Best Local Similarity:
   region of MHC
   Sequence 945
  Alignment Scores:
   20
  64
   40
   124
   181
  241
   301
  120
   361
  140
  421
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  Best Local Sin
Query Match:
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631
   240
  691
   260
  745
   280
  805
   300
   AAV12068;
                               220
   RESULT
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  The invention relates to a novel class II major histocompatibility complex (MHC) antigenic determinant of a mucous membrane invasive protein and the T-cell antigenic determinant of a mucous membrane invasive protein and the extracellular region of a class II MHC molecule or at least part of the extracellular region of the class II MHC molecule having an amino acid sequence comprising one or more deletions, substitutions or additions. The molecule of the invention may be useful for detecting an antigen-specific CD4+ T-cell by flow cytometry and for presenting a microorganism-derived mucous membrane invasive protein as an antigen. The method of the invention enables efficient detection of antigen-specific activation of CD4+ T-cells in the mucous membrane. The current sequence is that of the class II major histocompatibility complex-related I-bacibie coil, heat-labile toxin B subunit (LTB)-leucine
  108
  156
  210
  270
   119
  330
   GluTyrTrpAsnSerGlnProGluIleLeuGluArgThrArgAlaGluValAspThrAla 139
   390
  159
   450
  179
   510
   199
   570
   219
  59
   79
   66
  63
  LeuSerSerProGlyThrGluGlyGlyAsnSerlleCysPheSerProSerLeuGluHis 39
  ------GGTAAAGAGAA
  ProlleValValSerGlySerTrpAspGlyGlyGlyGlyGlySerLeuValProArgGlySer
  109 ATGGTTATCATT----ACATTTAAGGGTGGTGGTGGTTCTTTAGTTCCTAGA-----
  -----GGTGGTAGTGAAAGGCATTTCGTGTACCAGTTCATGGGCGAGTGCTACTTCACC
  ArgTyrAspSerAspValGlyGluTyrArgAlaValThrGluLeuGlyArgProAspAla
  CysArgHisAsnTyrGluGlyProGluThrSerThrSerLeuArgArgLeuGluGlnPro
  AsnValAlaIleSerLeuSerArgThrGluAlaLeuAsnHisHisAsnThrLeuValCys
  SerValThrAspPheTyrProAlaLysIleLysValArgTrpPheArgAsnGlyGlnGlu
   TCAGTGACAGATTTCTACCCAGCCAAGATCAAAGTGCGCTGGTTCCGGAATGGCCAGGAG
  1 MetAlaLeuGlnIlePro---SerLeuLeuLeuSerAlaAlaValValLeuMetVal
   GlyGlyGlySerGluArgHisPheValValGlnPheLysGlyGluCysTyrThr
  TGCAGACACAACTACGAGGGGCGGAGACCCACACCTCCCTGCGGCGGCTTGAACAGCCC
  GluThrvalGlyvalSerSerThrGlnLeulleArgAsnGlyAspTrpThrPheGlnVal
   AATGTCGTCATCTCCCTGTCCAGGACAGGCCCTCAACCACCACACACTCTGGTCTGC
   80 AsnGlyThrGlnArglleArgLeuValThrArgTyrlleTyrAsnArgGluGluTyrVal
   Sequence 915 BP; 228 A; 242 C; 271 G; 174 T; 0 U; 0 Other;
   915
242
116
32
114
5
  Length:
Matches:
Conservative:
Mismatches:
Indels:
  zipper (LZ)-BirA fusion cDNA of the invention.
  CTGAGCTCCCCACTGGCTTTGGCTGGAGACTCC---
                           Example 3; SEQ ID NO 13; 30pp; Japanese.
  Gaps:
   US-10-048-116B-6 (1-306) x ADQ31228 (1-915)
  7.14e-106
1219.00
84.9%
79.6%
75.2%
  Percent Similarity:
Best Local Similarity:
  .gnment Scores:
ğ
  331
  160
  20
   64
  40
   9
   157
  211
   100
   120
   451
  140
   391
  180
   511
   200
   Query Match:
DB:
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279
  This nucleotide sequence comprises a PCR product obtained by amplification of mouse splencoyte cDNA using primers (see AAV12065 and AAV12066) designed for the amplification of IAd beta chain full-length cDNA. IAd alpha chain cDNA (see AAV12067) has been similarly obtained. The IAd sequences were cloned into metallothionein promoter (see AAV12062)-driven vector pRmHa-3 prior to sequencing. Major histocompatibility complex (MHC) class II IAd heterodimers were expressed at the cell surface of transfected Drosophila Schneider 2 (ATCC CRL 10974) cells. The invention relates to the preparation and use of synthetic antigen presenting matrices, in particular antigen presenting cells such as insect cells that have been transfected to produce MHC
   259
   804
  299
   Major histocompatibility class II antigen; MHC class II; T cell; T lymphocyte; Th1; Th2; activation; CD4+; antigen presenting cell; APC; autoimmune disease; diabetes; multiple sclerosis; autoimmune thyroiditis; systemic lupus erythromatosus; myasthenia gravis; Crohn's disease; inflammatory bowel disease; allergy; asthma; contact sensitivity; immunotherapy; therapy; IAd beta chain; mouse; ds; circular; cyclic.
   S
   864
   Synthetic antigen presenting cell for activating CD4+ T cells - useful treat autoimmune disease, e.g. diabetes, multiple sclerosis, Crohn's disease and inflammatory bowel disease, or allergy, e.g. asthma and
   ProSerLeuLysSerProlleThrValGluTrpArgAlaGlnSerGluSerAlaArgSer
GAGACGGTGGGGGTCTCATCCACACACTTATTAGGAATGGGGACTGGACCTTCCAGGTC
                                    LeuValMetLeuGluMetThrProHisGlnGlyGluValTyrThrCysHisValGluHis
   CTGGTCATGCTGGAGATGACCCCTCGGGGGGGGAGGTCTACACCTGTCACGTGGAGCAT
   LysGlyGlyGlyGlySerThrThrAlaProSerAlaGlnLeuLysLysLysLeuGlnAla
   creerrecececedarecaeracaerecareaecreaerreaaaaaaaaaerecaeeca
  LeulyslyslyslandlaGlnLeulysTrpLysLeuGlnAlaLeulysLysLysLeuAla
   CTTAAGAAAAAGAACGCTCAGCTGAAGTGGAAACTTCAAGCCCTCAAGAAGAAACTCGCC
  PA;
  Peterson
  Jackson MR,
   Example 2; Page 94-96; 141pp; English.
  Karlsson L,
  AAV12068 standard; cDNA; 4724 BP
   97WO-US008697.
  96US-0018175P.
  Murine IAd beta chain cDNA.
   (first entry)
  GlnHisHisHis 303
  865 CAGCTGCATCAT 876
   (SCRI ) SCRIPPS RES INST
  Webb SR, Wingvist O,
   WPI; 1998-041895/04.
  contact sensitivity
  WO9746256-A1.
   Mus musculus
   22-MAY-1997;
  23-MAY-1996;
   08-JUN-1998
   11-DEC-1997.
```

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1068
   1008
   1128
antigen presenting molecules with one or more accessory molecules. The matrices are used to activate naive CD4+ T cells and to shift the ongoing activation state into a preferred differentiated population of Th1 or Th2 cells. Applications include the treatment of autoimmune disease, e.g. diabetes, multiple sclerosis, autoimmune thyroiditis, systemic lupus erythematosus, myasthenia gravis, Crohn's disease and inflammatory bowel disease, or an allergy, e.g. asthma and contact sensitivity
   240
  120
   160
   828
   180
   200
   948
  510
   537
   537
  588
   648
  708
   768
  40
  9
  80
   TyrAspSerAspValGlyGluTyrArgAlaValThrGluLeuGlyArgProAspAlaGlu
  ACAGTGGGGGTCTCATCCACACACGTTATTAGGAATGGGGACTGGACCTTCCAGGTCCTG
  SerSerProGlyThrGluGlyGlyAsnSerIleCysPheSerProSerLeuGluHisPro
  GlyGlyGlySerGluArgHisPheValValGlnPheLysGlyGluCysTyrTyrThrAsn
  GlyThrGlnArg1leArgLeuValThrArgTyr1leTyrAsnArgGluGluTyrValArg
   GGGACGCAGCGCATACGGCTCGTGACCAGATACATCTACAACCGGGAGGAGTACGTGGC
   TyrTrpAsnSerGlnProGluIleLeuGluArgThrArgAlaGluValAspThrAlaCys
  ArgHisAsnTyrGluGlyProGluThrSerThrSerLeuArgArgLeuGluGlnProAsn
   ValAlalleSerLeuSerArgThrGluAlaLeuAsnHisHisHisHeuValCysSer
  ThrValGlyValSerSerThrGlnLeuIleArgAsnGlyAspTrpThrPheGlnValLeu
   GTCATGCTGGAGATGACCCCTCATCAGGAGAGGTCTACACCTGCCATGTGGAGCATCCC
   AGCCTGAAGAGCCCCCATCACTGTGGAGTGGAGGCACAGTCCGAGTCTGCCCGGAGCAAG
   MetAlaLeuGlnIleProSerLeuLeuLeuSerAlaAlaValValValLeuMetValLeu
  41 IleValValSerGlySerTrpAspGlyGlyGlyGlySerLeuValProArgGlySerGly
  538 -----TCCGAAAGGCATTTCGTGGTCCAGTTCAAGGGCGAGTGCTACTACAAC
  TACTGGAACAGCCAGCCGGAGATCCTGGAGCGAACGCGGGGCCGAGGTGGACACGGCGTGC
   ValMetLeuGluMetThrProHisGlnGlyGluValTyrThrCysHisValGluHisPro
  SerieulysSerProlleThrValGluTrpArgAlaGlnSerGluSerAlaArgSerLys
  Sequence 4724 BP; 1196 A; 1194 C; 1200 G; 1134 T; 0 U; 0 Other;
   4724
228
1
1
37
   Length:
Matches:
Conservative:
Mismatches:
Indels:
   511 AGCAGCCCAGGGACTGAGGGCGGAAAC------
  Gaps:
   US-10-048-116B-6 (1-306) x AAV12068 (1-4724)
  1.85e-99
1161.50
85.8%
85.4%
  Percent Similarity:
Best Local Similarity:
Query Match:
DB:
   Alignment Scores:
Pred. No.:
  537
   21
  61
  81
   589
  101
  649
  121
  709
   141
  169
   161
   829
   889
   201
   949
   1009
  241
   181
  221
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/*tags e
/note= "binding site for primer #363 (see AAT04271)"
complement(177. .226)
  (see AAT04270)"
  /*tag= h
'note= "probable primer binding site (primer #270)"
31. .959
   cell;
  = "probable primer binding site (primer #271)".568
   (primer #272) "
  (primer #259)"
  #366} "

    18 /*tag= a
    10ce "probable primer binding site (primer #76)" complement(40. .74)

   /*tag= n
/note= "probable primer binding site (primer #59)"
  *tag= g
note= "primer #365 (see AAT04272) binding site"
85. .403
  Major histocompatability complex; MHC; T-cell receptor; TCR; autoimmune disease; immunodeficiency disease; immune response; immunoproliferation disease; graft-host rejection; therapy; B
  '*tag= m
'note= "probable primer binding site (primer
.000. .1013
  "IA beta chain beta 2 region"
  site
  primer #362
   Bite
  /note= "primer #364 binding site"
complement(212. .266)
  (NAJE-) NAT JEWISH CENT IMMUNOLOGY & RESPIRATORY
  "hybrid IA beta chain"
  note = "probable primer binding 42. .976
  primer binding
   *tag= b
note= "binding site for
   63. .143
/*tag= d
/note= "leader region"
complement(140. .191)
Location/Qualifiers
  /*tag= k
/note= "probable
323. .850
  BP.
  DNA; 1013
   94US-00207481
  /*tag= c
/product= '
  /product= '
  (first entry)
   3. .959
   M12.C3; pM12-IAb-Ea; 88.
   *tag=
   '+tag=
'note= '
   *tag=
  *tag=
   IA beta chain
  standard;
   04-MAR-1994;
  03-MAR-1995;
  16-APR-1996
  WO9523814-A1
  primer_bind
  primer_bind
  primer_bind
  sig_peptide
   primer_bind
   primer_bind
  primer_bind
   mat_peptide
  primer bind
   primer_bind
   primer_bind
   primer_bind
   primer_bind
   Synthetic
  AAT04269;
  AAT04269
  Hybrid
                                    RESULT
  ------GlyGlyGlyGly 264
```

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```
/label= I-Ad beta chain leader
/note= "murine MHC class II I-Ad gene beta chain leader
  MHC; major histocompatibility complex; PCR; polymerase chain reaction; T cell activity modulator; antagonist; immune disorder; allergy; multiple sclerosis; insulin-dependent diabetes mellitus; rheumatoid arthritis; myasthenia gravis; ds.
  ThrPheGlnValLeuValMetLeuGluMetThrProHisGlnGlyGluValTyrThrCys
HisValGluHisProSerLeuLysSerProlleThrValGluTrpArgAlaGlnSerGlu
  CACGTGGAGCATCCCAGCCTGAAGAGCCCCATCACTGTGGAGTGGAGTGGAGGCCACAGTCTGAG
  AsnGlyGlnGluGluThrValGlyValSerSerThrGlnLeuIleArgAsnGlyAspTrp
   ACCTTCCAGGTCCTGGTCATGCTGGAGATGACCCCTCGGCGGGGGAGAGGTCTAYACCTGT
  domain"
   domain'
   gene beta-2 domain"
   II I-Ad gene beta-1 domain'
  Vector SCE1-derived single chain gene encoding MHC fusion complex
   II I-Ad gene alpha-2
   II I-Ad gene alpha-2
   /label= OVA 323-339
/note= "chicken ovalbumin residues 323-339"
138. -167
/*tag= d
   /*tag= g
/note= "24 residue peptide linker"
307. .1067
   /*tag= d
/note= "10 regidue linker peptide"
168. .452
   SerAlaArgSerLys------GlyGlyGlyGly 264
   825 rcreccrecaccaacarcrreacceccarcecece
   /*tag= f
//label= I-Ad beta2
//ncte= "muxine MHC class II I-Ad
735. .806
  /note= "murine MHC class
   /*tag= h
/label= I-Ad alphal
/note= "murine MHC class
1068. 1352
  /*tag= i
|label= I-Ad_alpha2
|note= "murine MHC class
|353. .1379
  Location/Qualifiers 6. .1382
   /*tag= e
/label= I-Ad_betal
   /*tag= b
/label= I-Ad_beta_
  ВЪ
  /note= "EE tag
   AAT17588 standard; DNA; 1382
  sequence"
87. .137
  (first entry)
  ø
   *tag=
   /*tag=
  misc_feature
  26-SEP-1996
   misc_feature
   misc_feature
   misc_feature
  misc_feature
   misc_feature
  misc_feature
  misc_feature
   sig_peptide
   Synthetic.
   645
  216
                585
  196
   705
   236
  765
   AAT17588;
  256
  Key
  AAT17588
  RESULT
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   셤
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  요
  This sequence represents a hybrid IA beta chain gene. This sequence contains a fragment of the IE alpha chain (residues 56-73), as well as a linker and cleavage site. This sequence was transfected into a B cell line (M12.C3) using plasmid pM12-IAb-Ea. It was found that the encoded sequence was expressed in these cells. Complexes such as this may be used to regulate an immune response. The complexes are capable of being recognised by a TCR alone or in combination with additional MHC proteins. These complexes are useful for therapeutic purposes and experimental purposes. They can also be used as reagents for the treatment of diseases including autoimmune diseases, immunodeficiency diseases,
   227
  344
  115
   404
  135
   155
   175
   AGCAGCCCAGGGACTGAGGGCGGAGACTCC-----GAAGCTAGCTTGAGGCTCAG 173
  ---GGGGGAAGTGGAGGTGTGAAAAGCCATTTCGTGTACCAGTTCATGGCCAG
   464
  524
   584
   176 ThrLeuValCysSerValThrAspPheTyrProAlaLysIleLysValArgTrpPheArg 195
   Peptide-MHC complex comprising antigenic peptide, linker and MHC segment - useful as reagents for the treatment of diseases including auto-immune diseases, immuno-stimulatory diseases or graft-host rejection.
  SerSerProGlyThrGluGlyGlyAsnSerIleCysPheSerProSerLeuGluHisPro 40
   55
  75
   95
   56 ProArgGlySerGlyGlyGlyGlySerGluArgHisPheValValGlnPheLysGlyGlu
  CysTyrTyrThrAsnGlyThrGlnArgIleArgLeuValThrArgTyrIleTyrAsnArg
   GluGluTyrValArgTyrAspSerAspValGlyGluTyrArgAlaValThrGluLeuGly
   ArgProAspAlaGluTyrTrpAsnSerGlnProGluIleLeuGluArgThrArgAlaGlu
  ValAspThrAlaCysArgHisAsnTyrGluGlyProGluThrSerThrSerLeuArgArg
  465 GIGGACACGGIGIGCAGACACAACTACGAGGGCCGGAGACCCACACCTCCCTGCGGGGG
   LeuGluGlnProAsnValAlaIleSerLeuSerArgThrGluAlaLeuAsnHisHisAsn
   MetAlaLeuGlnIleProSerLeuLeuSerAlaAlaValValValLeumetValLeu
   ------ileValValSerGlySerTrpAspGlyGlyGlyGlySerLeuVal
  174 GGTGCACTGGCCAACATTGCTGTCGACAAGGCTGGAGGTGGATCCGGTGGA-----
  Sequence 1013 BP; 220 A; 272 C; 327 G; 192 T; 0 U; 2 Other;
  immunoproliferation diseases, and graft-host rejection
  Length:
Matches:
Conservative:
Mismatches:
  Indels:
   Gaps:
  US-10-048-116B-6 (1-306) x AAT04269 (1-1013)
  Example 2; Page 65; 94pp; English
  2.31e-99
1151.00
86.8%
84.6%
71.0%
            ä
            Marrack
  1995-320543/41.
   Percent Similarity:
Best Local Similarity:
  P-PSDB; AAR82538
   Alignment Scores:
           Kappler JW,
  21
   123
  96
   96
  345
   116
   63
   41
   228
  136
   156
   Query Match:
DB:
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180
   404
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   584
   644
   645 GTCATGCTGGAGATGACCCCTCATCAGGGAGGTCTACACCTGCCATGTGGAGCATCCC 704
   SerleulysSerProlleThrValGluTrpArgAlaGlnSerGluSerAlaArgSerLys 260
  Single chain major histocompatibility complex comprising linked alpha and
beta chains – useful for suppressing an immune response to an auto:immune
                  ValThrAspPheTyrProAlaLysIleLysValArgTrpPheArgAsnGlyGlnGluGlu
   ValMetLeuGluMetThrProHisGlnGlyGluValTyrThrCysHisValGluHisPro
  TyrTrpAsnSerGlnProGluIleLeuGluArgThrArgAlaGluValAspThrAlaCys
  ArgHisAsnTyrGluGlyProGluThrSerThrSerLeuArgArgLeuGluGlnProAsn
   AGACACAAACTACGAGGGGCGGAGAGACCAGCACCTCCCTGCGGCGGCTTGAACAGCCCAAT
   ValAlaIleSerLeuSerArgThrGluAlaLeuAanHisHisAsnThrLeuValCysSer
   GTCGCCATCTCCCTGTCCAGGACAGACCCCTCAACCACCACAACACTCTGGTCTGTTCG
  Thr ValGly Val Ser Ser Thr Gln Leu Ile Arg Asn Gly Asp Trp Thr Phe Gln Val Leu
   585 ACAGTGGGGGTCTCATCCACACAGCTTATTAGGAATGGGGACTGGACCTTCCAGGTCCTG
   Greacadartricracecadeceadareaagreecreerreagaaregeecadead
  Construction; major histocompatibility complex; MHC; fusion complex;
  Ä
  Wong
  Location/Qualifiers 6. .1382
  Burkhardt M,
   ВР.
   AAT86989 standard; DNA; 1382
  GCCGTGGTGGTTCC 770
  GlyGlyGlyGlySer 265
   97WO-US001617
  96US-00596387
   SCE1 single chain gene; ss.
  27-MAR-1998 (first entry)
   SCE1 single chain gene
   (DADE-) DADE INT INC
  WPI; 1997-402555/37.
  Rhode PR, Jiao J,
  P-PSDB; AAW29214.
   30-JAN-1997;
  31-JAN-1996;
  WO9728191-A1
   07-AUG-1997.
  261
   AAT86989;
  Synthetic
   345
  141
  405
   525
   756
   101
  121
   161
  465
  181
  201
   221
   241
  RESULT 8
  Key
  AAT86989
                             요
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   AAT17588 encodes a murine MHC fusion complex capable of modulating T cell activity encoded by the vector SCE1. The MHC fusion complex comprises at least one MHC molecule containing a peptide-binding groove and a presenting peptide covalently linked to the MHC molecule and opt. a presenting peptide covalently linked to the MHC molecule and opt. a transformate domain. DNA encoding a MHC fusion complex may be cloned into a host cell to express that encoding a MHC fusion complex or a single chain fusion complex by then molecule may be used to vaccinate a mammal against a targeted disorder. The fusion complexes may be used to suppress an immune response in an mimal suffering from an immune disorder e.g. multiple sclerosis, insulin-dependent diabetes mallitus, rheumatoid arthritis, mysathenia gravis or chronic allergies. The complexes may also be used in the treatment of livestock and pets such as cats and dogs. The MHC fusion complexes can be produced such that they contain a single antigenic peptide including one of known structure, additionally a wide range of peptides can be presented for T cell interaction
   66 AGCAGCCCAAGGACCTTAAGTATCTCTCAGGCTGTTCACGCTGCTCACGCTGAA----- 119
  GlyThrGlnArg1leArgLeuValThrArgTyrIleTyrAsnArgCluGluTyrValArg 100
  120 ATCAACGAAGCTGGTCGTGCTAGCGGAGGGGCGGAAGC---------GGCGGA 164
  165 GGGGGAAACTCCGAAAGGCATTTCGTGGTCCAGTTCAAGGGCGAGTGCTACTACACAAC 224
  9
  80
   20
  65
  SerSerProGlyThrGluGlyGlyAsnSerIleCysPheSerProSerLeuGluHisPro 40
   11eValValSerGlySerTrpAspGlyGlyGlyGlySerLeuValProArgGlySerGly
   GlyGlyGlySerGluArgHisPheValValGlnPheLysGlyGluCysTyrThrAsn
   MetAlaLeuGlnIleProSerLeuLeuLeuSerAlaAlaValValValLeuMetValLeu
  Major histocompatability complex fusion complex for modulating T celactivity - used in the treatment of immune disorders, e.g. multiple sclerosis, IDDM and rheumatoid arthritis.
   Sequence 1382 BP; 320 A; 374 C; 404 G; 284 T; 0 U; 0 Other;
  Edwards AC;
  Length:
Matches:
Conservative:
Mismatches:
Indels:
  Grammer S,
  Gaps:
  US-10-048-116B-6 (1-306) x AAT17588 (1-1382)
   Example 17; Fig 29; 210pp; English.
  Weidanz JA,
  95WO-US009816
   94US-00283302
95US-00382454
   1.3e-98
1145.00
87.2%
85.7%
70.7%
   Chavaillaz P, Jiao J;
   DADE-) DADE INT INC.
   Rhode PR,
  WPI; 1996-129343/13.
   Percent Similarity:
Best Local Similarity:
   P-PSDB; AAR98907
WO9604314-A1
  31-JUL-1995;
   29-JUL-1994;
  01-FEB-1995;
                               15-FEB-1996.
   Alignment Scores:
  Ä,
   н
  21
  41
   19
   8
  Query Match:
DB:
  Wong
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RESULT
   140
  119
   164
   224
   100
   284
   120
   344
   404
   160
  464
   180
   524
  200
  584
  220
  644
   704
  SerLeuLysSerProlleThrValGluTrpArgAlaGlnSerGluSerAlaArgSerLys 260
   -----ACTAGTGGTGGCGGTGGCAGC 755
   20
  65
  9
   80
  SerSerProGlyThrGluGlyGlyAsnSerIleCysPheSerProSerLeuGluHisPro 40
  AGCAGCCCAAGGACCTTAAGTATCTCTCAGGCTGTTCACGCTGCTCACGCTGAAA
   ATGCCTCTGCAGATCCCCAGCCTCCTCTCAGCTGCTGTGGTGGTGGTGTGTGATGGTGCTG
  41 jleValValSerGlySerTrpAspGlyGlyGlyGlyGlySerLeuValProArgGlySerGly
   ATCAACGAAGCTGGTCGTGCTAGCGGAGGGCGGAAGC---------GGCGGA
  GlyGlyGlySerGluArgHisPheValValGlnPheLysGlyGluCysTyrThrAsn
   GlyThrGlnArglleArgLeuValThrArgTyrIleTyrAsnArgGluGluTyrValArg
  GGGACGCAGCGCATACGGCTCGTGACCAGATACATCTACAACCGGGAGGAGTACGTGCGC
  TyrAspSerAspValGlyGluTyrArgAlaValThrGluLeuGlyArgProAspAlaGlu
  285 Taccacaccaccaccaccacaraccecceccacccaaccacaccaccaccacac
  TyrTrpAsnSerGlnProGluIleLeuGluArgThrArgAlaGluValAspThrAlaCys
  TACTGGAACAGCCAGCCGGAGATCCTGGAGCGAACGCGGGCCGAGGTGGACACGCGTGC
   141 ArgHisAsnTyrGluGlyProGluThrSerThrSerLeuArgArgLeuGluGlnProAsn
  AGACACAACTACGAGGGGCGGGAGACCAGCACCTCCCTGCGGCGGCTTGAACAGCCCAAT
  GTGGCCATCTCCCTGTCCAGGACAGAGGCCCTCAACCACCACAAAAACACTCTGGTCTGTTCG
  ValThrAspPheTyrProAlaLysIleLysValArgTrpPheArgAsnGlyGlnGluGlu
  GTGACAGATTTCTACCCAGCCAAGATCAAAGTGCGCTGGTTCAGGAATGGCCAGGAGGAG
   ThrValGlyValSerSerThrGlnLeuIleArgAsnGlyAspTrpThrPheGlnValLeu
   ACAGTGGGGGTCTCATCCACACACTTATTAGBAATGGGGACTGGACCTTCCAGGTCCTG
   ValMetLeuGluMetThrProHisGlnGlyGluValTyrThrCysHisValGluHisPro
  MetAlaLeuGlnIleProSerLeuLeuLeuSerAlaAlaValValValLeuMetValLeu
   GGGGGAAACTCCGAAAGGCATTTCGTGGTCCAGTTCAAGGGCGAGTGCTACTACACCAAC
  ValAlaIleSerLeuSerArgThrGluAlaLeuAsnHisHisAsnThrLeuValCysSer
   Greatscresasarsaccercareassasasasacreracecarstscassarce
 e.g. multiple sclerosis, rheumatoid arthritis, diabetes
  Sequence 1382 BP; 320 A; 373 C; 405 G; 284 T; 0 U; 0 Other;
  major
   the construction
   fusion complexes
  Conservative:
Mismatches:
  Length:
Matches:
   Indels:
                               Example 17; Page 140-141; 217pp; English
   AGCCTGAAGAGCCCCATCACTGTGGAGTGG
   US-10-048-116B-6 (1-306) x AAT86989 (1-1382)
  The present sequence was used in histocompatibility complex (MHC)
   GlyGlyGlyGlySer 265
   1.3e-98
1145.00
87.2%
85.7%
  Percent Similarity:
Best Local Similarity:
   gnment Scores:
   21
   120
  61
   165
   225
   345
  405
  525
   585
  221
   645
   261
   81
  101
  121
  161
   465
  181
   201
   241
  Query Match:
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The invention relates to a major histocompatibility complex (WHC) fusion complex (I) comprising an MHC molecule that contains a peptide-binding groove, and a presenting peptide covalently (e.g. an antigenic peptide) linked to the MHC molecule, where (I) is capable of modulating the complex, where the MHC molecule is a class II MHC (e.g. mouse I-Ad or I-As or human HIA-DRI (human leukocyte antigen-DRI)), a multivalent MHC than complex comprising two or more linked complexes, identifying a peptide that can modulate the activity of T cells (involving introducing into host cells cloning vectors that each contain the fusion complex DNA, culturing the host cells under conditions suitable for expression of the MHC fusion complex, and selecting host cells that express MHC fusion complex and selecting host cells that expression of the C expression vector comprising DNA that codes for the alpha and beta chains of the MHC fusion complex. The DNA constructs can contain beta chains of the fusion complex. Also included are inducing an immune response in a mammal (including vaccinating a mammal against a targeted disorder, by administering DNA sequence comprising a fusion complex, or DNA sequence comprising a single chain fusion molecule) and suppressing an immune response in a mammal (including vaccinating a mammal against a targeted disorder, by administering DNA sequence comprising a single chain fusion molecule) and suppressing an immune response in a mammal a DNA sequence comprising a transmembrane domain, and a public that contains a transmembrane domain, and a contain and contains a transmembrane domain, and a public that the contains a transmembrane domain, and a public that contains a transmembrane domain, and a public that contains a transmembrane domain, and a public that contains a transmembrane domain, and a public that contains a transmembrane domain, and a public that contains a transmembrane domain and a public that contains a transmembrane domain.
  presenting peptide that is a T cell receptor (TCR) antagonist or partial
   Novel major histocompatibility complex fusion complex having presenting peptide covalently linked to MHC molecule containing peptide-binding groove, used for suppressing immune response in multiple sclerosis,
  MHC; major histocompatibility complex; gene therapy; fusion complex; peptide-binding groove; T cell modulation; class II MHC; vaccine; autoximmune disorder; multiple sclerosis; rheumatoid arthritis; insulin-dependent diabetes mellitus; myasthenia gravis; immunogen; chronic allergy; mouse; ds; I-Ad; gene.
  Edwards AC;
   Mouse MHC I-Ad/Ova 323-339 synthetic gene SCE1.
  ŝ
   Grammer
  Example 17; Fig 29; 126pp; English.
  Weidanz JA,
   BP.
   94US-00283302.
95US-00382454.
97US-00776084.
   770
   ACA60744 standard; DNA; 1382
   06-JUL-2001; 2001US-00900379
   (first entry)
GGCGGTGGTGGTTCC
   Jiao JJJ;
  (DADE-) DADE INT INC.
  Rhode PR,
   WPI; 2003-341126/32.
P-PSDB; ABU72108.
   US2002198144-A1.
  Wong HC, Rhode
Chavaillaz P,
   29-JUL-1994;
01-FEB-1995;
17-JAN-1997;
   16-JUN-2003
   26-DEC-2002
   Mus sp.
Synthetic.
  allergies.
  ACA60744;
   756
```

```
705 AGCCTGAAGAGCCCCATCACTGTGGAGTGG----
   Rhode PR, Weidanz JA,
   770
   DNA; 1385
  95WO-US009816
   94US-00283302
95US-00382454
  (first entry)
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   *tag=
  (DADE-) DADE INT INC.
   AAT17586 standard;
   misc_feature
   misc feature
   misc_feature
  misc_feature
  misc_feature
  31-JUL-1995;
   29-JUL-1994;
01-FEB-1995;
  misc feature
   misc_feature
  misc_feature
   26-SEP-1996
  WO9604314-A1
   sig_peptide
   15-FEB-1996
   Synthetic.
   AAT17586;
   756
                          261
   Wong HC,
  RESULT 10
   Key
  AAT1758
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  셤
  agonist and is covalently linked to the MHC protein, or DNA sequence coding for the fusion complex which is a single chain fusion molecule. The methods are useful for identifying a peptide that can modulate the activity of T cells, inducing an immune response in a mammal (including vaccinating a mammal against a targeted disorder) and for suppressing an immune response in a mammal. The disorders include an autoimmune disorder such as multiple sclerosis, insulin-dependent disbetes mellitus, rheumatoid arthritis, myasthenia gravis or chronic allergies. The present sequence encodes a mouse MHC class II I-Ad fusion complex of the
  140
   119
  120
   344
  180
   200
  240
   100
   164
  224
   284
   404
  ArgHisAsnTyrGluGlyProGluThrSerThrSerLeuArgArgLeuGluGlnProAsn 160
   464
  524
  584
   644
   704
   SerLeuLysSerProlleThrValGluTrpArgAlaGlnSerGluSerAlaArgSerLys 260
   20
   65
  9
   80
  SerSerProGlyThrGluGlyGlyAsnSerIleCysPheSerProSerLeuGluHisPro 40
   AGCAGCCCAAGGACCTTAAGTATCTCTCAGGCTGTTCACGCTGCTCACGCTGAA-----
   ValAlaIleSerLeuSerArgThrGluAlaLeuAsnHisHisHisAsnThrLeuValCysSer
  ValThrAspPheTyrProAlaLys1leLysValArgTrpPheArgAsnGlyGluGluGlu
  ATGGCTCTGCAGATCCCCAGCCTCCTCTCAGCTGCTGTGGTGTGTGATGGTGCTG
   GlyGlyGlySerGluArgHisPheValValGlnPheLysGlyGluCysTyrTyrThrAsn
   Gregecarcrecergrecaggacagaecercraaceacacaacarcregrerger
  ThrvalGlyvalSerSerThrGlnLeuIleArgAsnGlyAspTrpThrPheGlnValLeu
   ValMetLeuGluMetThrProHisGlnGlyGluValTyrThrCysHisValGluHisPro
  GTCATGCTGGAGATGACCCCTCATCAGGAGAGGTCTACACCTGCCATGTGGAGCATCCC
   MetAlaLeuGlnIleProSerLeuLeuLeuSerAlaAlaValValValLeuMetValLeu
   IleValValSerGlySerTrpAspGlyGlyGlyGlySerLeuValProArgGlySerGly
  120 ATCAACGAAGCTGGTCGTGCTAGCGGAGGGGGGGGAAGC-------GGCGGA
  GGGGGAAACTCCGAAAGGCCATTTCGTGGTCCAGTTCAAGGCCGAGTGCTACTACACCAAC
   GlyThrGlnArg1leArgLeuValThrArgTyrIleTyrAsnArgGluGluTyrValArg
  GGGACGCAGCGCATACGGCTCGTGACCAGATACATCTACAACCGGGAGGAGTACGTGCGC
  TyrAspSerAspValGlyGluTyrArgAlaValThrGluLeuGlyArgProAspAlaGlu
  TyrTrpAsnSerGlnProGluIleLeuGluArgThrArgAlaGluValAspThrAlaCys
  GTGACAGATTTCTACCCAGCCAAGATCAAAGTGCGCTGGTTCAGGAATGGCCAGGAGGAG
  ACAGTGGGGGTCTCATCCACACACTTATTAGGAATGGGGACTGGACCTTCCAGGTCCTG
   Sequence 1382 BP; 320 A; 373 C; 405 G; 284 T; 0 U; 0 Other;
  1382
227
4
24
10
   Conservative:
Mismatches:
Indels:
Gaps:
  Length:
Matches:
   US-10-048-1168-6 (1-306) x ACA60744 (1-1382)
   1.3e-98
1145.00
87.2%
85.7%
  Percent Similarity:
Best Local Similarity:
   Alignment Scores:
  invention
  41
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   61
  165
  81
  225
   101
  285
   121
  345
   141
  405
   161
  465
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   645
  Best Local S
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---ACTAGTGGTGGCGGTGGCAGC 755
  beta chain leader
   T cell activity modulator; antagonist; polymerase chain reaction; multiple sclerosis; insulin-dependent diabetes mellitus; rheumatoid arthritis; myasthenia gravis; ds.
  gene alpha-2 domain"
   domain"
  domain"
  Vector SSC1-derived single chain gene encoding MHC fusion complex
   II I-Ad gene alpha-2
   beta-1
  beta-2
  *tag= c
/label= OVA_323-339
/note= "chicken ovalbumin residues 323-339"
  Grammer S, Edwards AC;
  gene
  gene
   gene
  *tag= g
note= "24 residue peptide linker"
07. .1067
  note= "10 residue linker peptide
   6..86
/*tag= b
/label= I-Ad_beta_chain_leader
/note= "murine MHC class II I-Ad e
   *tag= f
|label= I-Ad beta2
|note="murine MHC class II I-Ad |
  I-Ad
   II I-Ad
  Η
  *tag= h
|label= I-Ad_alphal
|note= "murine MHC class
| 1352
   /label= I-Ad_alpha2
/note= "murine MHC class
1353. .1382
   class
   Location/Qualifiers
   *tag= e
label= I-Ad betal
note= "murine MHC
   /*tag= j
/note= "6xHis tag*
   ВР
```

584

180 524 220

644

240

260

```
Single chain major histocompatibility complex comprising linked alpha and beta chains - useful for suppressing an immune response to an auto:immune disease. e-g- multiple sclerosis, rheumatoid arthritis, diabetes mellitus, etc.
   ThrValGlyValSerSerThrGlnLeuIleArgAsnGlyAspTrpThrPheGlnValLeu
  ValThrAspPheTyrProAlaLysIleLysValArgTrpPheArgAsnGlyGlnGluGlu
   585 ACAGTGGGGGTCTCATCCACACAGCTTATTAGGAATGGGGACTGGACCTTCCAGGTCCTG
                            465 GTCGCCATCTCCCTGTCCAGGACAGAGCCCTCAACCACCACAACACTCTGGTCGTCTCG
   ValMetLeuGluMetThrProHisGlnGlyGluValTyrThrCysHisValGluHisPro
       161 ValAlaIleSerLeuSerArgThrGluAlaLeuAsnHisHisAsnThrLeuValCysSer
  SerLeuLysSerProlleThrValGluTrpArgAlaGlnSerGluSerAlaArgSerLys
   major histocompatibility complex; MHC; fusion complex;
  Sequence 1385 BP; 316 A; 383 C; 399 G; 287 T; 0 U; 0 Other;
  The present sequence was used in the construction of major histocompatibility complex (MHC) fusion complexes
   1385
227
   Length:
Matches:
  Н
Н
   Example 17; Page 135-137; 217pp; English.
  Wong
  Location/Qualifiers
6. .1385
/*tag= a
  Rhode PR, Jiao J, Burkhardt M,
   BP.
   AAT86987 standard; DNA; 1385
   97WO-US001617.
  261 GlyGlyGlyGlySer 265
   756 Gecesirecrecirce 770
   96US-00596387
   1.3e-98
1145.00
   Construction; major histoco
SSC1 single chain gene; ss
  entry)
  SSC1 single chain gene.
  (DADE-) DADE INT INC
  WPI; 1997-402555/37.
  (first
  P-PSDB; AAW29212
   31-JAN-1996;
   30-JAN-1997;
  WO9728191-A1
  27-MAR-1998
   Alignment Scores:
  07-AUG-1997.
   Synthetic.
  201
  AAT86987;
   181
   221
  241
  ..
No...
   Key
   AAT86987
   à
   a
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  셤
  ò
  셤
  ò
   윱
  AAT17586 encodes a murine MHC fusion complex capable of modulating T cell activity encoded by the vector SSC1. The MHC fusion complex comprises at least one MHC molecule containing a peptide-binding groove and a presenting peptide covalently linked to the MHC molecule and opt. a presenting peptide covalently linked to the MHC molecule and opt. a transmembrane domain. DNA encoding a MHC fusion complex may be cloned into a host cell to express that modulate, pref. antagonise, T cell cetivity. DNA encoding a MHC fusion complex or a single chain fusion molecule may be used to vaccinate a mammal against a targeted disorder. The fusion complexes an immune response in an inmal suffering from an immune disorder e.g. multiple sclerosis, insulindependent diabetes mellitus, rheumatoid arthritis, mysathenia gravis or chronic allergies. The complexes and dogs. The MHC fusion complexes can be produced such that they contain a single antignic peptide including one of known structure, additionally a wide range of peptides can be presented for T cell interaction
   120
  140
   GlyThrGlnArgileArgLeuValThrArgTyrIleTyrAsnArgGluGluTyrValArg 100
  AGCAGCCCAAGGACCTTAAGTATCTCTCAGGCTGTTCACGCTGCTCACGCTGAA---- 119
  164
  224
   284
  404
   ArgHisAsnTyrGluGlyProGluThrSerThrSerLeuArgArgLeuGluGlnProAsn 160
   20
  6 ATGGCTCTGCAGATCCCCAGCCTCCTCTCAGCTGCTGTGGTGGTGCTGATGGTGCTG 65
   9
  80
   SerSerProGlyThrGluGlyGlyAsnSerlleCysPheSerProSerLeuGluHisPro 40
  Major histocompatability complex fusion complex for modulating T cell activity - used in the treatment of immune disorders, e.g. multiple sclerosis, IDDM and rheumatoid arthritis.
  TyrAspSerAspValGlyGluTyrArgAlaValThrGluLeuGlyArgProAspAlaGlu
  TACTGGAACAGCCAGCCGGAGATCCTGGAGCGAACGCGGGCCCAAGGTGGACACGGCGTGC
  1 MetAlaLeuGlnIleProSerLeuLeuLeuSerAlaAlaValValValLeuMetValLeu
  41 ileValValSerGlySerTrpAspGlyGlyGlyGlyGlySerLeuValProArgGlySerGly
  TyrTrpAsnSerGlnProGluIleLeuGluArgThrArgAlaGluValAspThrAlaCys
  Sequence 1385 BP; 316 A; 384 C; 398 G; 287 T; 0 U; 0 Other;
   1385
227
4
24
10
3
   Length:
Matches:
Conservative:
Mismatches:
Indels:
  US-10-048-116B-6 (1-306) x AAT17586 (1-1385)
   Example 17; Fig 27; 210pp; English.
   1.3e-98
1145.00
87.2$
85.7$
 Jiao J;
                                    WPI; 1996-129343/13.
   Percent Similarity:
Best Local Similarity:
Query Match:
  P-PSDB; AAR98905
Chavaillaz P,
  Alignment Scores:
  120
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119
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  TyrAspSerAspValGlyGluTyrArgAlaValThrGluLeuGlyArgProAspAlaGlu 120
  140
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   524
   200
  584
  220
  704
  SerieulysSerProlleThrValGluTrpArgAlaGlnSerGluSerAlaArgSerLys 260
   755
   ValMetLeuGluMetThrProHisGlnGlyGluValTyrThrCysHisValGluHisPro 240
  40
   9
   80
   SerSerProGlyThrGluGlyGlyAsnSerIleCysPheSerProSerLeuGluHisPro
   GlyThrGlnArgileArgLeuValThrArgTyrileTyrAsnArgGluGluTyrValArg
   TyrTrpAsnSerGlnProGluIleLeuGluArgThrArgAlaGluValAspThrAlaCys
   ArgHisAsnTyrGluGlyProGluThrSerThrSerTeleuhrgArgLeuGluGlnFroAsn
   ValThrAspPheTyrProAlaLysIleLysValArgTrpPheArgAsnGlyGlnGluGlu
   ThrvalGlyvalSerSerThrGlnLeuIleArgAsnGlyAspTrpThrPheGlnValLeu
  ATGGCTCTGCAGATCCCCAGCCTCCTCTCAGCTGCTGTGGTGGTGCTGATGGTGCTG
   GlyGlyGlySerGluArgHisPheValValGlnPheLysGlyGluCysTyrThrAsn
||||||
   Greacagarricracccaeccaagarcaaagrececresrrcaggaaresccaegagga
  GGGACGCAGCGCATACGGCTCGTGACCAGATACATCTACAACCGGGAGGAGTACGTGGC
  345 TACTGGAACAGCCAGCCGGAGATCCTGGAGCGAACGCGGGCCGAGGTGGACACGCCGGGCGTGC
  ValAlaIleSerLeuSerArgThrGluAlaLeuAsnHisHisAsnThrLeuValCysSer
   Greecearcreteresagacagacecercaaceceacacacacacreteres
  ACAGTGGGGGTCTCATCCACACACACACTTATTAGGAATGGGGACTGGACCTTCCAGGTCCTG
  MetalaLeuGlnIleProSerLeuLeuLeuSerAlaAlaValValValLeuMetValLeu
   AGACACAACTACGAGGGCCGGAGACCAGCACCTCCCTGCGGGCGTTGAACAGCCCAAT
  705 AGCCTGAAGAGCCCCATCACTGTGGAGTGG-----ACTAGTGGTGGCGGTGGCAGC
  41 IleValValSerGlySerTrpAspGlyGlyGlyGlySerteuValProArgGlySerGly
    3 10 4
3 0 1 6
 Conservative:
Mismatches:
Indels:
  Mouse MHC I-Ad/Ova 323-339 synthetic gene SSC1.
                                      Gaps:
   US-10-048-116B-6 (1-306) x AAT86987 (1-1385)
  ΒP
   ACA60742 standard; DNA; 1385
  GlyGlyGlyGlySer 265
   756 Gecedicarderice 770
  (first entry)
 87.2%
85.7%
70.7%
             Similarity:
Percent Similarity:
  16-JUN-2003
   99
   81
  21
   61
  101
  285
   405
  261
   121
   221
  141
  161
   465
   181
  201
            Best Local Si
Query Match:
DB:
   RESULT 12
   ACA60742
  g
   g
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   셤
  22225222
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```
The invention relates to a major histocompatibility complex (WHC) fusion complex (I) comprising an MHC molecule that contains a peptide-binding grove, and a presenting peptide covalently (e.g. an antigonic peptide) coveraged to the MHC molecule; where (I) is capable of modulating the cartivity of a T cell. Also included are a DNA construct coding for the activity of a T cell. Also included are a DNA construct coding for the complex, where the MHC molecule is a class II MHC (e.g. mouse I-Ad or I-C As, or human HLA-DR1 (human leukocyte antigen-DR1), a multivalent MHC (figure or more linked complexes, identifying a complex comprising two or more linked complexes, identifying a complex chan modulate the activity of T cells (involving introducing introducing the host cells under coditions sustable for expression of the complex that modulate the activity of T cells (involving introducing complex that modulate the activity of T cells), a single recombinant expression vector comprising DNA that codes for the alpha and beta chains of the MHC protein, a single recombinant expression vector comprising DNA that codes for a T cells), a single recombinant complex and beta chains of the MHC fusion complex. The DNA constructs can contain the terologous leader peptide sequences and Kozak sequence for efficient expression of the fusion complex. The DNA constructs can contain the esponse in a mammal (including vaccinating a mammal aby administering DNA sequence complex, the included are inducing an immune response in a mammal aby administering an expression vector, encoding a contain and a presenting peptide that is a T cell receptor (TCR) antagonist or partial agoint the fusion complex which is a single chain fusion and a presenting peptide that is a T cell receptor (TCR) antagonist or partial and suppressing an immune response in a mammal solution and suppressing an immune response in a mammal solution and suppressing an immune response in a mammal solution and suppressing an immune response in a mammal solution and suppressing
   Novel major histocompatibility complex fusion complex having presenting peptide covalently linked to MHC molecule containing peptide-binding groove, used for suppressing immune response in multiple sclerosis,
MHC; major histocompatibility complex; gene therapy; fusion comple
peptide-binding groove; T cell modulation; class II MHC; vaccine;
autoimmune disorder; multiple sclerosis; rheumatoid arthritis;
insulin-dependent diabetes mellitus; myasthenia gravis; immunogen;
  such as multiple sclerosis, insulin-dependent diabetes mellitus, rheumatoid arthritis, myasthenia gravis or chronic allergies. The sequence encodes a mouse MHC class II I-Ad fusion complex of the
   Edwards AC;
   s,
   Grammer
   chronic allergy; mouse; ds; I-Ad; gene
  Example 17; Fig 27; 126pp; English.
   Weidanz JA,
  95US-00382454.
  2001US-00900379
  94US-00283302
  Wong HC, Rhode PR, Weid
Chavaillaz P, Jiao JJJ;
   DADE-) DADE INT INC.
   2003-341126/32.
  P-PSDB; ABU72106.
   US2002198144-A1
  06-JUL-2001;
  29-JUL-1994;
  01-FEB-1995;
17-JAN-1997;
   26-DEC-2002
  Mus sp.
Synthetic.
   allergies
   nvention
```

287 T; 0 U; 0 Other;

ö

C; 399

Sequence 1385 BP; 316 A; 383

```
gene alpha-transmembrane
   chain leader
Colass II I-Ad gene beta chain leader
                                MHC; major histocompatibility complex; PCR; polymerase chain reaction; I cell activity modulator; antagonist; immune disorder; allergy; multiple sclerosis; insulin-dependent diabetes mellitus; rheumatoid arthritis; myasthenia gravis; ds.
  gene alpha-2 domain"
  gene alpha-2 domain"
  Major histocompatability complex fusion complex for modulating T cell activity - used in the treatment of immune disorders, e.g. multiple sclerosis, IDDM and rheumatoid arthritis.
   gene beta-1 domain"
   domain"
              SCT1-derived single chain gene encoding MHC fusion complex.
  gene beta-2
   /*tag= c
/label= OVA 323-339
/note= "chicken ovalbumin residues 323-339"
138. 167
   Edwards AC;
  *tag= g
note= "24 residue peptide linker"
  /*tag= d
/note= "10 residue linker peptide"
  /tags e
/label= I-Ad betal
/note= "murine MHC class II I-Ad
453. .734
  /*tag= f
/label= I.Ad_beta2
/note= "murine MHC class II I-Ad
735. .806
  /label= I-Ad alphal
/note= "murine MHC class II I-Ad
1068. .1352
  /*tag= i
hobel= I.Ad_alpha2
hoce="murine WHC class II I-Ad
1353. 1505
   /*tag= j
//abba= I-Ad_alpha-TM
/note= "murine MHC class II I-Ad
domain"
   Ś
   Grammer
  Location/Qualifiers
6. .1508
   'label= I-Ad beta c
'note= "murine MHC
   Example 17; Fig 28; 210pp; English.
   Weidanz JA,
   95WO-US009816.
   94US-00283302.
95US-00382454.
   .1067
  sequence"
   168. .452
   *tag= h
   *tag=
   *tag=
  Wong HC, Rhode PR, We
Chavaillaz P, Jiao J;
  (DADE-) DADE INT INC.
  WPI; 1996-129343/13.
   P-PSDB; AAR98906.
  misc_feature
  misc_feature
  misc_feature
   29-JUL-1994;
01-FEB-1995;
  misc_feature
   misc_feature
   misc_feature
  misc_feature
   misc_feature
  WO9604314-A1
  31-JUL-1995;
   sig peptide
  15-FEB-1996
  Synthetic
              Vector
   Key
   66 AGCAGCCCAAGGACCTTAAGTATCTCTCAGGCTGTTCACGCTGCTCACGCTGAA---- 119
  100
  TyrAspSerAspValGlyGluTyrArgAlaValThrGluLeuGlyArgProAspAlaGlu 120
   TyrTrpAsnSerGlnProGluIleLeuGluArgThrArgAlaGluValAspThrAlaCys 140
  224
  284
  ArgHisAsnTyrGluGlyProGluThrSerThrSerLeuArgArgLeuGluGlnProAsn 160
  ValThrAspPheTyrProAlaLyslleLysValArgTrpPheArgAsnGlyGlnGluGlu 200
   ThrValGlyValSerSerThrGlnLeuIleArgAsnGlyAspTrpThrPheGlnValLeu 220
  240
  464
  ValAlaileSerLeuSerArgThrGluAlaLeuAsnHisHisAsnThrLeuValCysSer 180
   524
  584
   585 ACAGTGGGGGTCTCATCCACACGTTATTAGGAATGGGGACTGGACCTTCCAGGTCCTG 644
  SerLeuLysSerProlleThrValGluTrpArgAlaGlnSerGluSerAlaArgSerLys 260
   755
   80
   20
   65
  9
   SerSerProGlyThrGluGlyGlyAsnSerIleCysPheSerProSerLeuGluHisPro 40
  GGGACGCACGCATACGCCTCGTGACCAGATACATCTACAACCGGGAGGAGTACGTGCGC
   MetAlaLeuGlnIleProSerLeuLeuLeuSerAlaAlaValValValLeuMetValLeu
  6 ArGGCTCTGCAGATCCCCAGCCTCCTCTCAGCTGCTGTGGTGGTGCTGATGGTGCTG
  GlyGlyGlySerCluArgHisPheValValGlnPheLysGlyGluCysTyrThrAsn
||||||
  GlyThrGlnArgileArgLeuValThrArgTyrIleTyrAsnArgGluGluTyrValArg
   41 jieValValSerGlySerTrpAspGlyGlyGlyGlySerLeuValProArgGlySerGly
  GTGACAGATTTCTACCCCAGCCAAGATCAAAGTGCGCTGGTTCAGGAATGGCCAGGAGGAG
   ValMetLeuGluMetThrProHisGlnGlyGluValTyrThrCysHisValGluHisPro
  1385
227
4
24
10
3
           Length:
Matches:
Conservative:
Mismatches:
  Gaps:
   (1-1385)
   BP.
   US-10-048-116B-6 (1-306) x ACA60742
  AAT17587 standard; DNA; 1508
   GlyGlyGlyGlySer 265
  770
           1.3e-98
1145.00
87.2%
85.7%
70.7%
   26-SEP-1996 (first entry)
  Gecernearectric
                         Percent Similarity:
Best Local Similarity:
Query Match:
DB:
Alignment Scores:
Pred. No.:
   63
  165
   225
   101
  141
   21
   81
  285
   121
  345
  405
  191
   465
   181
  525
   201
   221
   645
  241
   705
   261
   756
   AAT17587;
   RESULT 13
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```
AAT17587 encodes a murine MHC fusion complex capable of modulating T cell activity encoded by the vector SCT1. The MHC fusion complex comprises at least one MHC molecule containing a peptide-binding groove and a presenting peptide covalently linked to the MHC molecule and opt. a transmembrane domain. DNA encoding a MHC fusion complex may be cloned into a host cell to express the complex. The transformed cells may then be used to identify peptides that modulate, pref. antagonise, T cell activity. DNA encoding a MHC fusion complex or a single chain fusion molecule may be used to vaccinate a mammal against a targeted disorder. The fusion complexes may be used to suppress an immune response in an animal suffering from an immune disorder e.g. multiple sclerosis, insulin dependent diabetes mallitus, rheumacoid arthritis, mysathenia gravis or chronic allergies. The complexes may also be used in the treatment of livestock and pets such as cats and dogs. The MHC fusion complexes can be produced such that they contain a single antigenic peptide including one of known structure, additionally a wide range of peptides can be presented for T cell interaction
```

Sequence 1508 BP; 337 A; 414 C; 440 G; 317 T; 0 U; 0 Other;

```
164
   224
  GlyThrGlnArgileArgLeuValThrArgTyrileTyrAsnArgGluGluTyrValArg 100
   284
   101 TyraspSerAspValGlyGluTyrArgAlaValThrGluLeuGlyArgProAspAlaGlu 120
   140
  404
   141 ArgHisAsnTyrGluGlyProGluThrSerThrSerLeuArgArgLeuGluGlnProAsn 160
  464
   ValAlalleSerLeuSerArgThrGluAlaLeuAsnHisHisAsnThrLeuValCysSer 180
  524
  ValThrAspPheTyrProAlaLysIleLysValArgTrpPheArgAsnGlyGluGlu 200
   66 AGCAGCCCAAGGACCTTAAGTATCTCTCAGGCTGTTCACGCTGCTCACGCTGAA---- 119
  525 GTGACAGATTTCTACCCAGCCAAGATCAAAGTGCGCTGGTTCAGGAATGGCCAGGAGGG 584
   20
   65
   9
  80
   SerSerProGlyThrGluGlyGlyAsnSerIleCysPheSerProSerLeuGluHisPro 40
   6 ATGGCTCTGCAGATCCCCAGCCTCCTCTCAGCTGCTGTGGTGGTGGTGGTGGTGGTGCTG
  120 ATCAACGAAGCTGGTCGTGCTAGCGGAGGGGGGGAAGC--------GGCGGA
   165 GGGGGAAACTCCGAAAGGCATTTCGTGGTCCAGTTCAAGGGGGGAGTGCTACTACACCAAC
   225 GGGACGCATACGCTCGTGACCAGATACATCTACAACCGGGAGGAGTACGTGCGC
  TyrTrpAsnSerGlnProGluIleLeuGluArgThrArgAlaGluValAspThrAlaCys
  1 MetAlaLeuGlnIleProSerLeuLeuLeuSerAlaAlaValValValLeuMetValLeu
   GlyGlyGlySerGluArgHisPheValValGlnPheLysGlyGluCysTyrTyrThrAsn
  405 AGACACAACTACGAGGGCCGGAGACCAGCACCTCCCTGCGGGGGGCTTGAACAGCCCAAT
   GTCGCCATCTCCCTGTCCAGGACAGAGCCCTCAACCACCACCACAACACTCTGGTTCG
           Length:
Matches:
Conservative:
Mismatches:
Indels:
  US-10-048-116B-6 (1-306) x AAT17587 (1-1508)
          1.46e-98
1145.00
87.2%
85.7%
  Best Local Similarity:
Query Match:
                                       Percent Similarity:
Alignment Scores:
   21
   41
  121
  181
   61
   81
   465
  161
  셤
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```
704
                                      260
  Single chain major histocompatibility complex comprising linked alpha and beta chains - useful for suppressing an immune response to an auto:immune disease, e.g. multiple sclerosis, rheumatoid arthritis, diabetes mellitus, etc.
   20
  65
          SerLeuLysSerProlleThrValGluTrpArgAlaGlnSerGluSerAlaArgSerLys
   MetAlaLeuGlnIleProSerLeuLeuLeuSerAlaAlaValValValLeuMetValLeu
  major histocompatibility complex; MHC; fusion complex;
   BP; 337 A; 413 C; 441 G; 317 T; 0 U; 0 Other;
  The present sequence was used in the construction of major histocompatibility complex (MHC) fusion complexes
   1508
227
4
24
10
   Length:
Matches:
Conservative:
Mismatches:
Indels:
   Ä
  Example 17; Page 137-139; 217pp; English.
  Wong
  Gaps:
   US-10-048-116B-6 (1-306) x AAT86988 (1-1508)
   Location/Qualifiers
6. .1508
/*tag= a
   Σ
  BP.
   Rhode PR, Jiao J, Burkhardt
  261 GlyGlyGlyGlySer 265
   Gecelerefrectic 770
  AAT86988 standard; DNA; 1508
  97WO-US001617
   96US-00596387
  1.46e-98
1145.00
87.2%
85.7%
   Construction; major histoco
SCT1 single chain gene; ss.
   (first entry)
   SCT1 single chain gene.
   (DADE-) DADE INT INC.
  WPI; 1997-402555/37.
  Percent Similarity:
Best Local Similarity:
   P-PSDB; AAW29213
  Sequence 1508
   27-MAR-1998
   30-JAN-1997;
   31-JAN-1996;
  WO9728191-A1
   Alignment Scores:
Pred. No.:
   07-AUG-1997.
  Synthetic.
   756 (
221
                                       241
   AAT86988;
  Query Match:
DB:
   RESULT 14
   Key
  AAT86988
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```

complex class I complexes

HC; Wong

ט Jiao

Σ

Burkhardt

Acevedo J,

```
Single chain major histocompatibility
                            (SUNO-) SUNOL MOLECULAR CORP.
  1999-418411/35.
  P-PSDB; AAY27111
           29-OCT-1997;
  Rhode PR,
   345
   Query Match:
DB:
  8
  임
   ò
  임
  ò
   요
   à
  셤
  8
  셤
   8
   셤
   ò
   100
   120
  344
   140
  160
  220
  224
   284
  404
   180
  200
  164
   464
  524
  584
   644
  SerLeuLysSerProlleThrValGluTrpArgAlaGlnSerGluSerAlaArgSerLys 260
   9
  80
  Major histocompatibility complex; MHC; single chain MHC; sc-MHC; Ig; pebtide binding groove; immunoglobulin; T cell receptor; immune response;
  TACGACAGGACGTGGGGCGAGTACCGCGCGGTGACCGAGCTGGGGCGGCCAGACGCCGAG
SerSerProGlyThrGluGlyGlyAsnSerIleCysPheSerProSerLeuGluHisPro
                 66 AGCAGCCCAAGGACCTTAAGTATCTCTCAGGCTGTTCACGCTGCTCACGCTGAA----
   GlyGlyGlySerGluArgHisPheValValGlnPheLysGlyGluCysTyrTyrThrAsn
  GlyThrGlnArgileArgleuValThrArgTyrIleTyrAsnArgGluGluTyrValArg
  GGGACGCAGCGCTCGTCGTCGCCAGATACATCTACAACCGGGAGGAGTACGTGCGC
  TyrAspSerAspValGlyGluTyrArgAlaValThrGluLeuGlyArgProAspAlaGlu
  TyrTrpAsnSerGlnProGluIleLeuGluArgThrArgAlaGluValAspThrAlaCys
   345 TACTGGAACAGCCGGGAGTCTGGAGCGAACGGGGGCCGAGGTGGACACGGGGTGG
   ArgHisAsnTyrGluGlyProGluThrSerThrSerLeuArgArgLeuGluGlnProAsn
   ValThrAspPheTyrProAlaLys1leLysValArgTrpPheArgAsnGlyGlnGluGlu
   525 GTGACAGATTTCTACCCAGCCAAGATCAAAGTGCGCTGGTTCAGGAATGGCCAGGAGGAG
  ACAGTGGGGGTCTCATCCACACACTTATTAGGAATGGGGACTGGACTTCCAGGTCCTG
   ValAlaIleSerLeuSerArgThrGluAlaLeuAsnHisHisAsnThrLeuValCysSer
  ThrValGlyValSerSerThrGlnLeuIleArgAsnGlyAspTrpThrPheGlnValLeu
                                   IleValValSerGlySerTrpAspGlyGlyGlyGlySerLeuValProArgGlySerGly
  peptide binding groove; immunoglobulin; T cell receptor; immune
immune-related disorder; antigenic peptide; fusion protein; ss.
   Single chain IAd/OVA 323-229 MHC fusion protein encoding DNA.
  120 ATCAACGAAGCTGGTCGTGCTAGCGGAGGGGGGGGGAAGC-----
   BP
   AAX89069 standard; DNA; 1508
  GlyGlyGlyGlySer 265
   98WO-US021520
  (first entry)
   deceereereerrec
  14-SEP-1999
   13-OCT-1998;
   W09921572-A1
   06-MAY-1999
  Synthetic.
   61
   165
  81
  225
  585
                                    41
  101
  285
   121
   141
  405
  191
   465
   221
  181
  201
  645
  241
  261
   AAX89069;
  705
  RESULT
                셤
   쉱
  q
   셤
                                   ઠે
   ò
   엄
   ò
   ઠે
  셤
   ે છે
  셤
  ò
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  셤
   쉽
   g
   ò
  ઠે
   ઠે
   염
  ò
   g
  셤
  ò
```

```
The invention relates to new single chain major histocompatibility complex (sc-MHC) class II complexes that comprise a peptide binding groove, and a modified class II beta 2 chain or covalantly linked circulary for defection and analysis of peptide ligands, pathogenic Teclis, for functional, cellular and molecular assays. They can be used to identify and isolate T cell receptor and/or MHC agonists and antagonists. They can be used in vivo to compete with pathogenic antigen presenting cells involved in immune-related disorders. They can also be used to craise antibodies and to screen immune cells. It is also use in a method of suppressing an immune response in mammals. The sc-MHC complexes of suppressing an immune response II mammals. The sc-MHC complexes comprising modified class II beta 2 chains and/or Ig-C1 regions are soluble and provide enhanced yield. These MHC complexes also can contain single antigenic peptides readily isolated from expressing cells in significant quantities. The polyspecific MHC complexes also provide a means to detect cells expressing multiple target structures with a single complex. The present sequence represents a DNA encoding a single chain
  120
  100
  284
   164
   224
   404
  65
  9
   21 SerSerProGlyThrGluGlyGlyAsnSerIleCysPheSerProSerLeuGluHisPro 40
   80
   6 ATGGCTCTGCAGATCCCCAGCCTCCTCTCAGCTGCTGTGGTGGTGGTGGTGGTGGTGCTGT
   AGCAGCCCAAGGACCTTAAGTATCTCTCAGGCTGTTCACGCTGCTCACGCTGAA-----
  GlyGlyGlySerGluArgHisPheValValGlnPheLysGlyGluCysTyrTyrThrAsn
   GlyThrGlnArglleArgLeuValThrArgTyrIleTyrAsnArgGluGluTyrValArg
  TyrAspSerAspValGlyGluTyrArgAlaValThrGluLeuGlyArgProAspAlaGlu
   MetAlaLeuGlnIleProSerLeuLeuLeuSerAlaAlaValValValLeuMetValLeu
  41 ileValValSerGlySerTrpAspGlyGlyGlyGlySerLeuValProArgGlySerGly
   120 ATCAACGAAGCTGGTCGTGCTAGCGGAGGCGGGAAGC--------GGCGGA
  GGGGGAAACTCCGAAAGGCATTTCGTGGTCCAGTTCAAGGGCGAGTGCTACTACACCAAC
  GGGACGCGGGGGTACGGCTCGTGACCAGATACATCTACAACCGGGAGGAGTACGTGCGC
  Sequence 1508 BP; 337 A; 413 C; 441 G; 317 T; 0 U; 0 Other;
   Length:
Matches:
Conservative:
Mismatches:
Indels:
Gaps:
   (1-1508)
Example 1; Fig 1; 148pp; English
  US-10-048-116B-6 (1-306) x AAX89069
  1.46e-98
1145.00
87.2%
85.7%
   Percent Similarity:
Best Local Similarity:
   Alignment Scores:
   99
  165
  225
  285
   61
  81
   101
  121
```

| ò      | 141 ArgHisbanTyrGluGlyProGluThrSerThrSerLeuArgArgLeuGluGlnProAsn 160   |   |
|--------|------------------------------------------------------------------------|---|
| 엄      | 405 AGACACAACTACGAGGGGCGGGAGACCAGCACCTCCCTGCGGGGGCTTGAACAGCCCAAT 464   |   |
| È      | 161 ValAlalleSerLeuSerArgThrGluAlaLeuAsnHisHisHsAsnThrLeuValCysSer 180 | , |
| 셤      | 465 GTGGCCATCTCCCTGTCCAGGACAGAGGCCCTCAACCACCACAACACTCTGGTCTGTTGG 524   |   |
| ò      | 181 ValThrAspPheTyrProAlaLysIeLysValArgTrpPheArgAsnGlyGluGlu 200       |   |
| a      | 525 GTGACAGATTTCTACCCAGCCAAGATCGCCTGCTTCAGGATGGCCAGGAGGG 584           |   |
| È      | 201 ThrValGlyValSerSerThrGlnLeuIleArgAsnGlyAspTrpThrPheGlnValLeu 220   |   |
| q      | 585 ACAGTGGGGGTCTCATCCACAGCTTATTAGGAATGGGGACTGGACCTTCCAGGTCCTG 644     |   |
| ò      | 221 ValMetLeuGluMetThrProHisGlnGlyGluValTyrThrCysHisValGluHisPro 240   |   |
| ą      | 645 GTCATGCAGAGATGACCCCTCATCAGGAGAGGTCTACACCTGCCATGTGGAGCATCCC 704     |   |
| ò      | 241 SerLeuLysSerProlleThrValGluTrpArgAlaGlnSerGluSerAlaArgSerLys 260   |   |
| d<br>d | 105 AGCCTGAAGAGCCCCATCACTGGAGTGGACTAGTGGTGGCGGTGGCAGC 755              |   |
| ò      | 261 GlyGlyGlyGer 265                                                   |   |
| q      | 756 GGCGGTGGTTCC 770                                                   |   |
|        |                                                                        | _ |

Search completed: June 30, 2006, 01:48:08 Job time: 539.723 8ecs

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AR033964 Sequence
AR175097 Sequence
AX032545 Sequence
AX032545 Sequence
AX032543 Sequence
AX032543 Sequence
AR175095 Sequence
AR175095 Sequence
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AR033963 Sequence
AR152030 Sequence
AR15203 Sequence
AX032544 Sequence
AX032544 Sequence
AX032544 Sequence
AX032544 Sequence
AX03254 Sequence
AX03254 Sequence
AX03254 Mus muscu
AF15202 Mus muscu
AF253060 Mus muscu
AF253060 Mus muscu
AF19251 Mus muscu
AF253061 Mus muscu
AF19251 Mus muscu
AF19251 Mus muscu
AF19251 Mus muscu
AF19251 Mus muscu
AF19251 Mus muscu
AF19250 Mus muscu
AF19250 Mus muscu
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Patent: WO 0109194-A 2 08-EEB-2001;
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Unclassified.

K Kappler,J.W. and Marzack,P.
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Reppler, J.W. and Marrack, P.
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1 (bases 1 to 4724)

Webb, S.R., Wingvist, O., Karlsson, L., Jackson, M.R. and Peterson, P.A.
WHC class II antigen-presenting systems and methods for activating CD4+ T cells
Patent: US 6355479-A 8 12-MAR-2002;

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Rhode, P.R., Jiao, J.-A., Burkhardt, M. and Wong, H.C. Single chain MMC complexes and uses thereof Patent: US 5869270-A 123 09-FEB-1999;
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PAT 17-DEC-2001

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1 (Bases I to 1305)
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Patent: EP 0991477-A 13 03-MAY-2000;
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AL Patent: US 6232445-A 24 15-MAY-2001;
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Rhode, P.R., Jiao, J.-A., Burkhardt, M. and Wong, H.C. Single chain MHC complexes and uses thereof Patent: US 5869270-A 122 09-FEB-1999;
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JW Unclassified.
E 1 (bases 1 to 1508)

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